

Supplemental Material

Pesticide Exposure and Depression among Male Private Pesticide Applicators in the Agricultural Health Study

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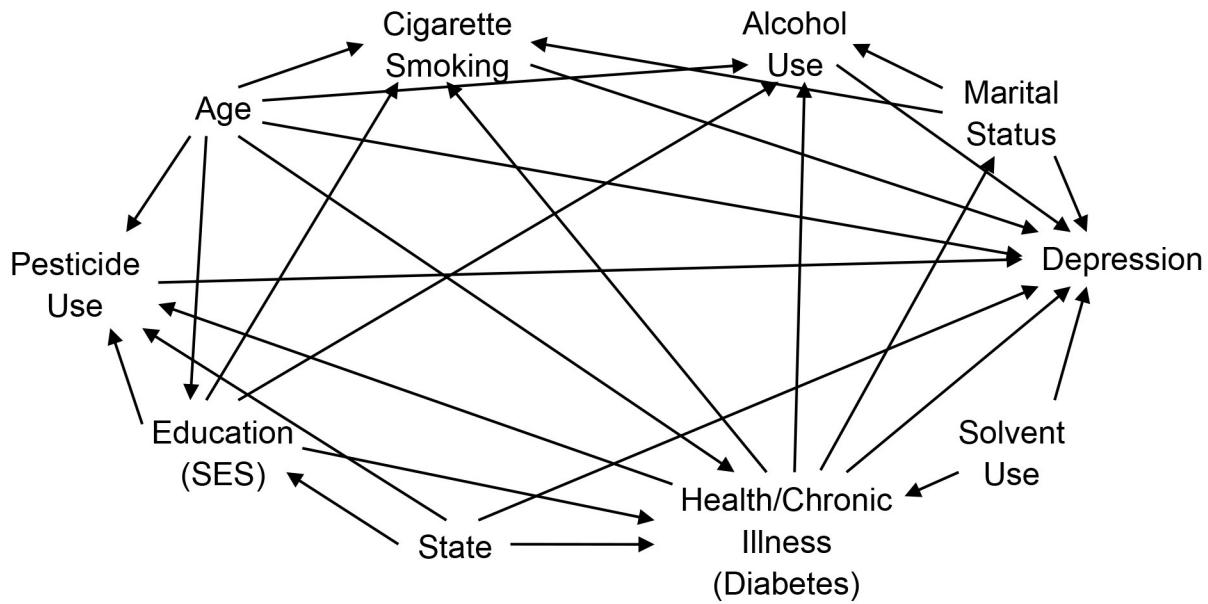


Figure S1. Directed acyclic graph to identify potential confounders that needed to be controlled to estimate the total effect of pesticide use on self-reported depression among male private pesticide applicators in the Agricultural Health Study. Note that switching the direction of the arrows from “Health/Chronic Illness (Diabetes)” to “Cigarette Smoking”, from “Health/Chronic Illness (Diabetes)” to “Alcohol Use”, and from “Health/Chronic Illness (Diabetes)” to “Marital Status” or adding an arrow from “Age” to “Marital Status” all result in the same two minimally sufficient adjustment sets as the directed acyclic graph shown here.

Methods, statistical analyses

We used standard formulas (Muller and Fetterman 2002; Stokes et al. 2000) to estimate linear, logistic, or ordinal logistic regression models to calculate the various stabilized weights (Hernán et al. 2000; Robins et al. 2000). In all models, the quantities of interest were the predicted probabilities of exposure, not missing covariate data, completing the farmer questionnaire, or not dropping out of the cohort. The populations to which the weights applied, types of models used, outcome of each model, and covariates included in each model were as follows:

Exposure/confounding weights

Population: 45,827 male applicators not missing data on depression at enrollment and at follow-up and not missing covariate data (for analyses involving information from the farmer questionnaire, this number was 20,471)

Model type: linear, logistic, or ordinal logistic (depending on the nature of the exposure variable)

Outcome: exposure (continuous, dichotomous, or ordered categories)

Covariates in models used to estimate numerators: intercept only

Covariates in models used to estimate denominators: age, diabetes, education, state

Missing covariate (diabetes and education) data weights

Population: 49,142 male applicators not missing data on depression at enrollment and at follow-up (for analyses involving information from the farmer questionnaire, this numbers was 22,300)

Model type: logistic

Outcome: not missing covariate data (dichotomous)

Covariates in models used to estimate numerators: exposure

Covariates in models used to estimate denominators: age, state, exposure, age*exposure, state*exposure

Missing farmer questionnaire weights (if appropriate)

Population: 45,827 male applicators not missing data on depression at enrollment and at follow-up and not missing covariate data

Model type: logistic

Outcome: completed farmer questionnaire (dichotomous)

Covariates in models used to estimate numerators: intercept only

Covariates in models used to estimate denominators: age, diabetes, education, state

Selection/drop out weights

Population: 45,827 male applicators not missing data on depression at enrollment and at follow-up and not missing covariate data (for analyses involving information from the farmer questionnaire, this number was 20,471)

Model type: logistic

Outcome: did not drop out of cohort (dichotomous)

Covariates in models used to estimate numerators: exposure

Covariates in models used to estimate denominators: age, diabetes, education, state, exposure, age*exposure, diabetes*exposure, education*exposure, state*exposure

For more detailed information on inverse probability weighting, see Cole and Hernán (2008), Hernán et al. (2000, 2004), Hernán and Robins (2006), Robins (1998), Robins et al. (2000), and Sato and Matsuyama (2003).

Table S1. More characteristics of male private pesticide applicators in the Agricultural Health Study.

| Characteristic | Non-cases [n (%)] | PRE-E ^a Cases [n (%)] | PRE-E ^a Adjusted OR ^b (95% CI) | PRE-B ^a Cases [n (%)] | PRE-B ^a Adjusted OR ^b (95% CI) | POST ^a Cases [n (%)] | POST ^a Adjusted OR ^b (95% CI) | p for difference among ORs ^c |
|--|----------------------|--|--|--|--|---------------------------------------|---|--|
| Total | 19,506 (100) | 474 (100) | | 540 (100) | | 688 (100) | | |
| Marital status | | | | | | | | |
| Married or living as married | 17,121 (88) | 402 (85) | Reference | 453 (84) | Reference | 620 (90) | Reference | |
| Divorced, separated, or widowed | 688 (4) | 39 (8) | 2.5 (1.8, 3.5) | 47 (9) | 2.7 (2.0, 3.6) | 22 (3) | 0.9 (0.6, 1.3) | |
| Never married | 1,676 (9) | 33 (7) | 1.3 (0.9, 2.0) | 38 (7) | 1.3 (0.9, 1.9) | 46 (7) | 0.7 (0.5, 1.0) | < 0.01 |
| Missing | 21 | 0 | | 2 | | 0 | | |
| Number of children in family | | | | | | | | |
| 0 | 2,787 (14) | 53 (11) | Reference | 67 (12) | Reference | 86 (13) | Reference | |
| 1-3 | 12,659 (65) | 309 (66) | 0.9 (0.7, 1.2) | 364 (68) | 0.8 (0.6, 1.1) | 478 (70) | 1.2 (1.0, 1.6) | |
| 4-5 | 3,248 (17) | 88 (19) | 0.9 (0.6, 1.3) | 83 (15) | 0.7 (0.5, 0.9) | 106 (15) | 1.2 (0.9, 1.6) | |
| ≥ 5 | 702 (4) | 18 (4) | 0.8 (0.4, 1.4) | 23 (4) | 0.8 (0.5, 1.4) | 17 (2) | 1.0 (0.6, 1.7) | 0.26 |
| Missing | 110 | 6 | | 3 | | 1 | | |
| Frequency of alcohol consumption during past year (times a week) | | | | | | | | |
| Never | 6,448 (33) | 172 (36) | Reference | 189 (35) | Reference | 219 (32) | Reference | |
| < 1 | 6,628 (34) | 162 (34) | 1.0 (0.8, 1.2) | 194 (36) | 1.0 (0.8, 1.3) | 245 (36) | 1.1 (0.9, 1.3) | |
| 1-4 | 5,281 (27) | 105 (22) | 0.8 (0.6, 1.1) | 132 (25) | 0.9 (0.7, 1.1) | 171 (25) | 0.9 (0.7, 1.1) | |
| > 4 | 1,073 (6) | 34 (7) | 1.2 (0.8, 1.8) | 22 (4) | 0.7 (0.4, 1.1) | 45 (7) | 1.2 (0.9, 1.7) | 0.45 |
| Missing | 76 | 1 | | 3 | | 8 | | |
| Cigarette smoking status | | | | | | | | |
| Never | 11,155 (57) | 230 (49) | Reference | 268 (50) | Reference | 336 (49) | Reference | |
| Past | 5,935 (31) | 175 (37) | 1.2 (1.0, 1.5) | 201 (37) | 1.3 (1.0, 1.5) | 236 (35) | 1.5 (1.2, 1.7) | |
| Current | 2,342 (12) | 66 (14) | 1.3 (1.0, 1.8) | 69 (13) | 1.2 (0.9, 1.6) | 112 (16) | 1.6 (1.2, 1.9) | 0.50 |
| Missing | 74 | 3 | | 2 | | 4 | | |
| Times visited medical doctor or medical assistant about health concern in past year^d | | | | | | | | |
| 0 | 3,887 (36) | 49 (16) | Reference | 43 (14) | Reference | 90 (24) | Reference | |
| 1 | 3,556 (33) | 72 (24) | 1.6 (1.1, 2.3) | 75 (24) | 1.9 (1.3, 2.8) | 125 (34) | 1.6 (1.2, 2.1) | |
| > 1 | 3,417 (31) | 183 (60) | 4.1 (3.0, 5.7) | 189 (62) | 5.3 (3.8, 7.5) | 154 (42) | 2.2 (1.7, 2.9) | < 0.01 |
| Missing | 130 | 2 | | 8 | | 2 | | |

| Characteristic | Non-cases [n (%)] | PRE-E ^a Cases [n (%)] | PRE-E ^a Adjusted OR ^b (95% CI) | PRE-B ^a Cases [n (%)] | PRE-B ^a Adjusted OR ^b (95% CI) | POST ^a Cases [n (%)] | POST ^a Adjusted OR ^b (95% CI) | p for difference among ORs ^c |
|---|----------------------|--|--|--|--|---------------------------------------|---|--|
| Years lived or worked on farm over lifetime^d | | | | | | | | |
| < 21 | 967 (9) | 17 (6) | 0.8 (0.5, 1.3) | 27 (9) | 1.3 (0.8, 2.0) | 41 (11) | 1.4 (0.9, 1.9) | |
| 21-30 | 1,387 (13) | 29 (10) | 1.0 (0.6, 1.4) | 39 (13) | 1.3 (1.0, 1.8) | 58 (16) | 1.4 (1.0, 1.9) | |
| > 30 | 8,466 (78) | 258 (85) | Reference | 243 (79) | Reference | 265 (73) | Reference | 0.32 |
| Missing | 170 | 2 | | 6 | | 7 | | |
| Size of farm worked last year (acres) | | | | | | | | |
| Didn't work on a farm or none | 591 (3) | 16 (3) | Reference | 21 (4) | Reference | 22 (3) | Reference | |
| < 50 | 2,415 (13) | 72 (16) | 1.1 (0.7, 2.0) | 77 (15) | 0.9 (0.6, 1.5) | 81 (13) | 0.8 (0.5, 1.4) | |
| 50-499 | 8,949 (48) | 228 (50) | 0.9 (0.5, 1.6) | 260 (50) | 0.7 (0.4, 1.2) | 301 (47) | 0.9 (0.6, 1.4) | |
| > 499 | 6,781 (36) | 144 (31) | 0.8 (0.5, 1.4) | 163 (31) | 0.6 (0.4, 1.0) | 238 (37) | 0.9 (0.6, 1.5) | 0.26 |
| Missing | 770 | 14 | | 19 | | 46 | | |
| Wear chemical resistant gloves when personally handle pesticides | | | | | | | | |
| No | 5,156 (26) | 129 (27) | Reference | 147 (27) | Reference | 198 (29) | Reference | |
| Yes | 14,350 (74) | 345 (73) | 1.0 (0.8, 1.2) | 393 (73) | 0.9 (0.8, 1.1) | 490 (71) | 0.9 (0.7, 1.0) | 0.68 |
| Ever have job off farm^d | | | | | | | | |
| No | 3,725 (34) | 91 (30) | Reference | 100 (32) | Reference | 110 (30) | Reference | |
| Yes | 7,151 (66) | 215 (70) | 1.2 (1.0, 1.6) | 211 (68) | 1.1 (0.9, 1.4) | 259 (70) | 1.2 (0.9, 1.5) | 0.83 |
| Missing | 114 | 0 | | 4 | | 2 | | |
| Exposed to solvents (other than gasoline) at longest held non-farm job^d | | | | | | | | |
| No | 8,961 (82) | 235 (77) | Reference | 247 (79) | Reference | 311 (84) | Reference | |
| Yes | 1,915 (18) | 71 (23) | 1.5 (1.1, 1.9) | 64 (21) | 1.2 (0.9, 1.6) | 58 (16) | 0.8 (0.6, 1.1) | 0.01 |
| Missing | 114 | 0 | | 4 | | 2 | | |

Abbreviations: CI, confidence interval; OR, odds ratio; POST, post-enrollment; PRE-B, pre-enrollment both; PRE-E, pre-enrollment enrollment only.

^aCases divided into three groups based on when the physician-diagnosis of depression occurred (before or after enrollment) and on when it was reported via the AHS contacts (at enrollment, at follow-up, or both). The PRE-E group included applicators who reported a previous diagnosis of

depression at enrollment, but who did not confirm their pre-enrollment diagnosis at follow-up. The PRE-B group included applicators who reported a previous diagnosis of depression at both enrollment and follow-up, or who reported a previous diagnosis at follow-up only but with an age at diagnosis less than their age at enrollment. The POST group included applicators who reported a previous diagnosis of depression at follow-up but not enrollment, and whose reported age at diagnosis equaled or exceeded their age at enrollment.^b Adjusted for age at enrollment (modeled with a cubic polynomial) and state of residence.^c Differences among case-group-specific ORs tested via Wald χ^2 tests.^d Data available only for 11,982 applicators who completed the farmer questionnaire.

Table S2. Summary statistics for the truncated (at the 1st and 99th percentiles) overall stabilized inverse probability weights^a for models in Table 2.

| Variable | Mean | Standard Deviation | 1st Percentile | 5th Percentile | Median | 95th Percentile | 99th Percentile |
|---|------|--------------------|----------------|----------------|--------|-----------------|-----------------|
| Cumulative lifetime days personally mixed or applied pesticides | | | | | | | |
| Categorical ^b | 1.05 | 0.32 | 0.58 | 0.67 | 0.97 | 1.68 | 2.27 |
| Trend ^c | 1.00 | 0.23 | 0.67 | 0.73 | 0.95 | 1.46 | 1.81 |
| Ever diagnosed with pesticide poisoning ^d | 0.99 | 0.34 | 0.52 | 0.57 | 0.95 | 1.61 | 2.42 |
| Ever experienced an incident of unusually high personal pesticide exposure ^d | 1.05 | 0.32 | 0.58 | 0.67 | 0.97 | 1.68 | 2.27 |

^aWeights adjusted for age at enrollment (modeled with a restricted, quadratic spline with knots at ages 40, 48, and 57 years based on percentiles of the age distribution in cases), ever diagnosed with diabetes, education level, state of residence, not missing covariate data (conditional on age, state, the exposure, and pairwise interaction terms between each covariate and the exposure), and not dropping out of the AHS cohort (conditional on age, diabetes, education, state, the exposure, and pairwise interaction terms between each covariate and the exposure).

^bCategory boundaries set at quartiles of cumulative lifetime days of pesticide use among all male private pesticide applicators. ^cUsed within-category medians. ^dData available only for 11,982 applicators who completed the farmer questionnaire. Weights additionally adjusted for completing the farmer questionnaire (conditional on age, diabetes, education, and state).

Table S3. Summary statistics for the truncated (at the 1st and 99th percentiles) overall stabilized inverse probability weights^a for models in Table 3.

| Ever personally mixed or applied | Mean | Standard deviation | 1st Percentile | 5th Percentile | Median | 95th Percentile | 99th Percentile |
|---|-------------|---------------------------|-----------------------|-----------------------|---------------|------------------------|------------------------|
| Fumigants | 0.99 | 0.50 | 0.34 | 0.42 | 0.81 | 2.00 | 2.70 |
| Aluminum phosphide | 0.99 | 0.21 | 0.60 | 0.75 | 0.96 | 1.41 | 1.75 |
| Carbon tetrachloride/carbon disulfide (80/20 mix) | 0.99 | 0.23 | 0.33 | 0.73 | 0.95 | 1.41 | 1.91 |
| Ethylene dibromide | 0.99 | 0.23 | 0.44 | 0.73 | 0.95 | 1.46 | 1.84 |
| Methyl bromide | 0.98 | 0.70 | 0.27 | 0.31 | 0.79 | 2.06 | 5.70 |
| Fungicides | 1.00 | 0.59 | 0.41 | 0.48 | 0.77 | 2.48 | 2.87 |
| Benomyl ^b | 0.99 | 0.41 | 0.25 | 0.42 | 0.88 | 1.63 | 3.28 |
| Captan | 0.99 | 0.22 | 0.56 | 0.72 | 0.96 | 1.43 | 1.85 |
| Chlorothalonil | 0.98 | 0.35 | 0.30 | 0.51 | 0.90 | 1.63 | 2.57 |
| Maneb/mancozeb | 0.98 | 0.41 | 0.24 | 0.36 | 0.88 | 1.63 | 3.07 |
| Metalaxyl | 0.99 | 0.54 | 0.35 | 0.41 | 0.79 | 2.18 | 2.75 |
| Ziram | 1.00 | 0.21 | 0.66 | 0.74 | 0.95 | 1.42 | 1.73 |
| Herbicides | 0.99 | 0.24 | 0.64 | 0.72 | 0.94 | 1.47 | 1.90 |
| Alachlor | 0.99 | 0.28 | 0.62 | 0.64 | 0.96 | 1.52 | 2.28 |
| Butylate | 0.98 | 0.30 | 0.53 | 0.57 | 0.97 | 1.44 | 2.60 |
| Chlorimuron-ethyl | 0.99 | 0.28 | 0.61 | 0.66 | 0.92 | 1.54 | 2.18 |
| Dicamba | 0.98 | 0.69 | 0.47 | 0.51 | 0.73 | 2.06 | 4.82 |
| EPTC | 0.98 | 0.38 | 0.51 | 0.55 | 0.94 | 1.33 | 3.60 |
| Glyphosate | 0.99 | 0.32 | 0.59 | 0.62 | 0.93 | 1.63 | 2.40 |
| Imazethapyr | 0.98 | 0.79 | 0.48 | 0.55 | 0.76 | 1.72 | 6.17 |
| Metolachlor | 0.99 | 0.35 | 0.61 | 0.62 | 0.91 | 1.63 | 2.59 |
| Paraquat | 0.99 | 0.43 | 0.36 | 0.47 | 0.85 | 1.85 | 2.38 |
| Pendimethalin | 1.00 | 0.22 | 0.63 | 0.73 | 0.96 | 1.44 | 1.79 |
| Petroleum oil | 0.99 | 0.28 | 0.62 | 0.63 | 0.96 | 1.47 | 2.34 |
| Trifluralin | 0.99 | 0.43 | 0.56 | 0.58 | 0.85 | 1.80 | 3.08 |
| Phenoxy herbicides | 0.99 | 0.34 | 0.58 | 0.61 | 0.90 | 1.70 | 2.34 |
| 2,4-D | 0.99 | 0.36 | 0.56 | 0.60 | 0.88 | 1.75 | 2.44 |
| 2,4,5-T | 0.98 | 0.35 | 0.39 | 0.55 | 0.92 | 1.58 | 2.86 |
| 2,4,5-TP | 0.99 | 0.23 | 0.44 | 0.72 | 0.95 | 1.41 | 1.94 |

| Ever personally mixed or applied | Mean | Standard deviation | 1st Percentile | 5th Percentile | Median | 95th Percentile | 99th Percentile |
|---|-------------|---------------------------|-----------------------|-----------------------|---------------|------------------------|------------------------|
| <i>Triazine herbicides</i> | 0.99 | 0.40 | 0.45 | 0.58 | 0.85 | 1.85 | 2.45 |
| Atrazine | 0.99 | 0.37 | 0.47 | 0.64 | 0.86 | 1.71 | 2.46 |
| Cyanazine | 0.98 | 0.59 | 0.51 | 0.55 | 0.80 | 1.52 | 4.33 |
| Metribuzin | 0.98 | 0.45 | 0.54 | 0.55 | 0.86 | 1.68 | 3.31 |
| Insecticides | 1.00 | 0.24 | 0.63 | 0.72 | 0.94 | 1.48 | 1.90 |
| <i>Carbamates^b</i> | 1.00 | 0.56 | 0.57 | 0.62 | 0.82 | 1.99 | 4.05 |
| Aldicarb | 0.99 | 0.46 | 0.29 | 0.40 | 0.88 | 1.77 | 3.38 |
| Carbaryl | 0.99 | 0.56 | 0.52 | 0.59 | 0.76 | 2.28 | 3.58 |
| Carbofuran | 0.99 | 0.26 | 0.57 | 0.65 | 0.93 | 1.47 | 2.18 |
| <i>Organochlorine insecticides</i> | 0.98 | 0.61 | 0.50 | 0.54 | 0.73 | 2.24 | 4.00 |
| Aldrin | 0.95 | 0.42 | 0.31 | 0.39 | 0.90 | 1.58 | 3.33 |
| Chlordane | 0.98 | 0.43 | 0.37 | 0.48 | 0.85 | 1.80 | 3.12 |
| DDT | 0.96 | 0.67 | 0.33 | 0.38 | 0.79 | 2.16 | 4.77 |
| Dieldrin | 0.97 | 0.26 | 0.25 | 0.52 | 0.95 | 1.40 | 2.04 |
| Heptachlor | 0.95 | 0.37 | 0.28 | 0.37 | 0.91 | 1.54 | 2.75 |
| Lindane | 0.99 | 0.26 | 0.48 | 0.57 | 0.94 | 1.40 | 2.37 |
| Toxaphene | 0.98 | 0.32 | 0.36 | 0.51 | 0.91 | 1.55 | 2.43 |
| <i>Organophosphate insecticides</i> | 0.99 | 0.27 | 0.58 | 0.70 | 0.92 | 1.55 | 2.03 |
| Chlorpyrifos | 0.99 | 0.23 | 0.67 | 0.69 | 0.95 | 1.44 | 1.95 |
| Coumaphos | 0.99 | 0.22 | 0.59 | 0.75 | 0.95 | 1.42 | 1.82 |
| Diazinon | 1.00 | 0.39 | 0.44 | 0.55 | 0.85 | 1.74 | 2.41 |
| Dichlorvos | 0.98 | 0.24 | 0.42 | 0.64 | 0.94 | 1.37 | 2.12 |
| Fonofos | 0.98 | 0.43 | 0.51 | 0.56 | 0.96 | 1.32 | 4.05 |
| Malathion | 0.99 | 0.28 | 0.60 | 0.67 | 0.91 | 1.58 | 2.13 |
| Parathion | 0.99 | 0.34 | 0.34 | 0.54 | 0.90 | 1.62 | 2.40 |
| Phorate | 0.99 | 0.43 | 0.52 | 0.58 | 0.97 | 1.48 | 3.59 |
| Terbufos | 0.99 | 0.38 | 0.59 | 0.61 | 0.93 | 1.68 | 2.90 |
| Trichlorfon | 1.00 | 0.21 | 0.66 | 0.74 | 0.95 | 1.43 | 1.78 |
| <i>Pyrethroid insecticides</i> | 0.99 | 0.27 | 0.55 | 0.60 | 0.95 | 1.44 | 2.24 |
| Permethrin (for animals) | 0.98 | 0.28 | 0.44 | 0.54 | 0.95 | 1.37 | 2.47 |
| Permethrin (for crops) | 0.99 | 0.24 | 0.58 | 0.70 | 0.95 | 1.47 | 1.86 |

Abbreviations: 2,4-D, (2,4-dichlorophenoxy)acetic acid; 2,4,5-T, (2,4,5-trichlorophenoxy)acetic acid; 2,4,5-TP, (*RS*)-2-(2,4,5-trichlorophenoxy)propionic acid; DDT, 1,1,1-trichloro-2,2-bis(4-chlorophenyl)ethane; EPTC, S-ethyl dipropyl(thiocarbamate).

^aWeights adjusted for age at enrollment (modeled with a restricted, quadratic spline with knots at ages 40, 48, and 57 years based on percentiles of the age distribution in cases), ever diagnosed with diabetes, education level, state of residence, not missing covariate data (conditional on age, state, the exposure, and pairwise interaction terms between each covariate and the exposure), and not dropping out of the AHS cohort (conditional on age, diabetes, education, state, the exposure, and pairwise interaction terms between each covariate and the exposure). ^bBenomyl is also included in carbamates.

Table S4. Summary statistics for the truncated (at the 1st and 99th percentiles) overall stabilized inverse probability weights^a for models in Table S5.

| Cumulative lifetime days personally mixed or applied ^{b,c} | Mean | Standard deviation | 1st Percentile | 5th Percentile | Median | 95th Percentile | 99th Percentile |
|---|------|--------------------|----------------|----------------|--------|-----------------|-----------------|
| Fumigants | | | | | | | |
| Aluminum phosphide ^d | | | | | | | |
| Categorical ^e | 0.99 | 0.31 | 0.53 | 0.63 | 0.93 | 1.63 | 2.12 |
| Trend ^f | 1.00 | 0.31 | 0.56 | 0.64 | 0.92 | 1.64 | 2.11 |
| Carbon tetrachloride/carbon disulfide (80/20 mix) ^d | | | | | | | |
| Categorical | 0.99 | 0.31 | 0.35 | 0.65 | 0.92 | 1.61 | 2.13 |
| Trend ^f | 1.00 | 0.31 | 0.57 | 0.64 | 0.93 | 1.63 | 2.12 |
| Ethylene dibromide ^d | | | | | | | |
| Categorical ^e | 1.00 | 0.35 | 0.50 | 0.62 | 0.92 | 1.72 | 2.35 |
| Trend ^f | 1.00 | 0.31 | 0.56 | 0.64 | 0.92 | 1.62 | 2.13 |
| Methyl bromide | | | | | | | |
| Categorical | 1.02 | 0.44 | 0.27 | 0.63 | 0.89 | 1.82 | 3.41 |
| Trend ^f | 1.00 | 0.21 | 0.67 | 0.74 | 0.95 | 1.43 | 1.76 |
| Fungicides | | | | | | | |
| Benomyl ^{d,g} | | | | | | | |
| Categorical ^e | 1.00 | 0.39 | 0.28 | 0.60 | 0.90 | 1.79 | 2.48 |
| Trend ^f | 1.00 | 0.31 | 0.55 | 0.64 | 0.92 | 1.63 | 2.12 |
| Captan | | | | | | | |
| Categorical | 1.00 | 0.22 | 0.58 | 0.73 | 0.96 | 1.41 | 1.81 |
| Trend ^f | 1.00 | 0.21 | 0.67 | 0.74 | 0.95 | 1.42 | 1.75 |
| Chlorothalonil | | | | | | | |
| Categorical | 1.00 | 0.28 | 0.44 | 0.70 | 0.94 | 1.56 | 2.16 |
| Trend ^f | 1.00 | 0.21 | 0.67 | 0.74 | 0.95 | 1.42 | 1.75 |
| Maneb/mancozeb ^d | | | | | | | |
| Categorical | 1.00 | 0.38 | 0.28 | 0.60 | 0.90 | 1.77 | 2.42 |
| Trend ^f | 1.00 | 0.31 | 0.57 | 0.64 | 0.92 | 1.62 | 2.12 |

| Cumulative lifetime days personally mixed or applied ^{b,c} | Mean | Standard deviation | 1st Percentile | 5th Percentile | Median | 95th Percentile | 99th Percentile |
|---|------|--------------------|----------------|----------------|--------|-----------------|-----------------|
| Metalaxyd | | | | | | | |
| Categorical | 1.08 | 0.58 | 0.34 | 0.55 | 0.90 | 2.38 | 3.41 |
| Trend ^f | 0.99 | 0.29 | 0.56 | 0.65 | 0.93 | 1.59 | 2.09 |
| Herbicides | | | | | | | |
| Alachlor | | | | | | | |
| Categorical | 1.00 | 0.23 | 0.60 | 0.67 | 0.96 | 1.42 | 2.01 |
| Trend ^f | 1.00 | 0.22 | 0.68 | 0.74 | 0.95 | 1.44 | 1.75 |
| Butylate ^d | | | | | | | |
| Categorical | 0.99 | 0.29 | 0.44 | 0.64 | 0.97 | 1.53 | 2.19 |
| Trend ^f | 1.00 | 0.31 | 0.57 | 0.64 | 0.93 | 1.63 | 2.12 |
| Chlorimuron-ethyl ^d | | | | | | | |
| Categorical | 0.99 | 0.33 | 0.56 | 0.63 | 0.92 | 1.68 | 2.24 |
| Trend ^f | 1.00 | 0.31 | 0.57 | 0.64 | 0.92 | 1.63 | 2.10 |
| Dicamba | | | | | | | |
| Categorical | 1.07 | 0.61 | 0.48 | 0.53 | 0.99 | 1.64 | 5.07 |
| Trend ^f | 1.00 | 0.21 | 0.69 | 0.74 | 0.96 | 1.41 | 1.75 |
| EPTC | | | | | | | |
| Categorical | 0.98 | 0.22 | 0.51 | 0.68 | 0.96 | 1.35 | 1.96 |
| Trend ^f | 1.00 | 0.21 | 0.67 | 0.74 | 0.95 | 1.42 | 1.75 |
| Glyphosate | | | | | | | |
| Categorical | 1.05 | 0.36 | 0.57 | 0.66 | 0.95 | 1.80 | 2.38 |
| Trend ^f | 1.00 | 0.22 | 0.67 | 0.74 | 0.95 | 1.44 | 1.77 |
| Imazethapyr | | | | | | | |
| Categorical | 1.04 | 0.60 | 0.51 | 0.55 | 0.94 | 1.70 | 5.79 |
| Trend ^f | 1.00 | 0.21 | 0.67 | 0.74 | 0.95 | 1.42 | 1.77 |
| Metolachlor | | | | | | | |
| Categorical | 1.00 | 0.27 | 0.59 | 0.66 | 0.95 | 1.45 | 2.29 |
| Trend ^f | 1.00 | 0.22 | 0.67 | 0.74 | 0.95 | 1.43 | 1.79 |
| Paraquat ^d | | | | | | | |
| Categorical | 1.04 | 0.46 | 0.33 | 0.58 | 0.90 | 2.04 | 2.74 |
| Trend ^f | 1.00 | 0.31 | 0.55 | 0.64 | 0.92 | 1.63 | 2.08 |

| Cumulative lifetime days personally mixed or applied^{b,c} | Mean | Standard deviation | 1st Percentile | 5th Percentile | Median | 95th Percentile | 99th Percentile |
|---|-------------|---------------------------|-----------------------|-----------------------|---------------|------------------------|------------------------|
| Pendimethalin ^d | | | | | | | |
| Categorical | 1.02 | 0.36 | 0.52 | 0.61 | 0.94 | 1.76 | 2.38 |
| Trend ^f | 0.99 | 0.31 | 0.57 | 0.64 | 0.92 | 1.63 | 2.13 |
| Petroleum oil ^d | | | | | | | |
| Categorical | 0.99 | 0.31 | 0.54 | 0.64 | 0.94 | 1.60 | 2.18 |
| Trend ^f | 1.00 | 0.31 | 0.57 | 0.63 | 0.93 | 1.65 | 2.14 |
| Trifluralin | | | | | | | |
| Categorical | 1.05 | 0.32 | 0.52 | 0.64 | 1.02 | 1.46 | 2.68 |
| Trend ^f | 1.00 | 0.21 | 0.68 | 0.74 | 0.95 | 1.42 | 1.78 |
| <i>Phenoxy herbicides</i> | | | | | | | |
| 2,4-D | | | | | | | |
| Categorical | 1.10 | 0.37 | 0.53 | 0.64 | 1.04 | 1.71 | 2.86 |
| Trend ^f | 1.00 | 0.21 | 0.68 | 0.74 | 0.95 | 1.42 | 1.76 |
| 2,4,5-T ^d | | | | | | | |
| Categorical | 0.99 | 0.33 | 0.33 | 0.55 | 0.92 | 1.63 | 2.41 |
| Trend ^f | 1.00 | 0.31 | 0.58 | 0.63 | 0.93 | 1.64 | 2.13 |
| 2,4,5-TP ^d | | | | | | | |
| Categorical ^e | 0.99 | 0.31 | 0.50 | 0.64 | 0.92 | 1.63 | 2.15 |
| Trend ^f | 1.00 | 0.31 | 0.57 | 0.63 | 0.93 | 1.63 | 2.13 |
| <i>Triazine herbicides</i> | | | | | | | |
| Atrazine | | | | | | | |
| Categorical | 1.05 | 0.28 | 0.54 | 0.66 | 1.02 | 1.51 | 2.26 |
| Trend ^f | 1.00 | 0.22 | 0.68 | 0.74 | 0.95 | 1.45 | 1.76 |
| Cyanazine | | | | | | | |
| Categorical | 1.02 | 0.40 | 0.51 | 0.58 | 0.98 | 1.41 | 3.65 |
| Trend ^f | 1.00 | 0.21 | 0.68 | 0.74 | 0.95 | 1.42 | 1.85 |
| Metribuzin ^d | | | | | | | |
| Categorical | 1.00 | 0.41 | 0.41 | 0.51 | 0.94 | 1.51 | 3.44 |
| Trend ^f | 1.00 | 0.30 | 0.57 | 0.63 | 0.93 | 1.62 | 2.08 |

| Cumulative lifetime days personally mixed or applied ^{b,c} | Mean | Standard deviation | 1st Percentile | 5th Percentile | Median | 95th Percentile | 99th Percentile |
|---|------|--------------------|----------------|----------------|--------|-----------------|-----------------|
| Insecticides | | | | | | | |
| Carbamates ^g | | | | | | | |
| Aldicarb ^d | | | | | | | |
| Categorical ^e | 1.00 | 0.43 | 0.33 | 0.59 | 0.89 | 1.88 | 2.92 |
| Trend ^f | 0.99 | 0.30 | 0.56 | 0.64 | 0.92 | 1.61 | 2.08 |
| Carbaryl ^d | | | | | | | |
| Categorical | 1.28 | 0.89 | 0.44 | 0.56 | 0.87 | 3.25 | 4.59 |
| Trend ^f | 1.00 | 0.31 | 0.55 | 0.64 | 0.92 | 1.63 | 2.15 |
| Carbofuran | | | | | | | |
| Categorical | 1.00 | 0.22 | 0.60 | 0.72 | 0.95 | 1.43 | 1.91 |
| Trend ^f | 1.00 | 0.21 | 0.68 | 0.74 | 0.95 | 1.42 | 1.74 |
| <i>Organochlorine insecticides</i> | | | | | | | |
| Aldrin ^d | | | | | | | |
| Categorical | 0.98 | 0.32 | 0.24 | 0.41 | 0.93 | 1.55 | 2.36 |
| Trend ^f | 1.00 | 0.30 | 0.57 | 0.64 | 0.92 | 1.62 | 2.08 |
| Chlordane ^d | | | | | | | |
| Categorical | 0.99 | 0.40 | 0.28 | 0.53 | 0.88 | 1.73 | 2.79 |
| Trend ^f | 1.00 | 0.31 | 0.56 | 0.64 | 0.93 | 1.63 | 2.09 |
| DDT ^d | | | | | | | |
| Categorical | 0.99 | 0.43 | 0.27 | 0.43 | 0.88 | 1.80 | 3.03 |
| Trend ^f | 1.00 | 0.31 | 0.56 | 0.64 | 0.93 | 1.65 | 2.12 |
| Dieldrin ^d | | | | | | | |
| Categorical ^e | 0.99 | 0.31 | 0.28 | 0.65 | 0.92 | 1.61 | 2.12 |
| Trend ^f | 1.00 | 0.31 | 0.57 | 0.63 | 0.93 | 1.64 | 2.11 |
| Heptachlor ^d | | | | | | | |
| Categorical | 0.98 | 0.32 | 0.25 | 0.56 | 0.92 | 1.57 | 2.26 |
| Trend ^f | 1.00 | 0.30 | 0.57 | 0.64 | 0.92 | 1.62 | 2.09 |
| Lindane ^d | | | | | | | |
| Categorical | 0.99 | 0.30 | 0.39 | 0.66 | 0.93 | 1.60 | 2.18 |
| Trend ^f | 1.00 | 0.31 | 0.56 | 0.64 | 0.92 | 1.63 | 2.09 |

| Cumulative lifetime days personally mixed or applied^{b,c} | Mean | Standard deviation | 1st Percentile | 5th Percentile | Median | 95th Percentile | 99th Percentile |
|---|-------------|---------------------------|-----------------------|-----------------------|---------------|------------------------|------------------------|
| Toxaphene ^d | | | | | | | |
| Categorical | 1.00 | 0.34 | 0.47 | 0.64 | 0.91 | 1.66 | 2.26 |
| Trend ^f | 1.00 | 0.31 | 0.55 | 0.63 | 0.93 | 1.63 | 2.12 |
| <i>Organophosphate insecticides</i> | | | | | | | |
| Chlorpyrifos | | | | | | | |
| Categorical | 1.00 | 0.22 | 0.66 | 0.73 | 0.95 | 1.45 | 1.84 |
| Trend ^f | 1.00 | 0.22 | 0.67 | 0.73 | 0.95 | 1.44 | 1.80 |
| Coumaphos | | | | | | | |
| Categorical | 1.00 | 0.21 | 0.65 | 0.75 | 0.96 | 1.41 | 1.76 |
| Trend ^f | 1.00 | 0.21 | 0.67 | 0.74 | 0.95 | 1.42 | 1.75 |
| Diazinon ^d | | | | | | | |
| Categorical | 1.02 | 0.39 | 0.37 | 0.62 | 0.91 | 1.80 | 2.44 |
| Trend ^f | 1.00 | 0.31 | 0.56 | 0.63 | 0.92 | 1.62 | 2.11 |
| Dichlorvos | | | | | | | |
| Categorical | 0.99 | 0.20 | 0.44 | 0.76 | 0.95 | 1.37 | 1.81 |
| Trend ^f | 1.00 | 0.21 | 0.67 | 0.74 | 0.95 | 1.42 | 1.75 |
| Fonofos | | | | | | | |
| Categorical | 0.98 | 0.22 | 0.53 | 0.65 | 0.96 | 1.34 | 1.90 |
| Trend ^f | 1.00 | 0.21 | 0.68 | 0.74 | 0.96 | 1.42 | 1.78 |
| Malathion ^d | | | | | | | |
| Categorical | 1.02 | 0.33 | 0.49 | 0.64 | 0.94 | 1.65 | 2.31 |
| Trend ^f | 1.00 | 0.31 | 0.56 | 0.64 | 0.92 | 1.63 | 2.10 |
| Parathion ^d | | | | | | | |
| Categorical | 1.00 | 0.35 | 0.41 | 0.62 | 0.91 | 1.72 | 2.31 |
| Trend ^f | 1.00 | 0.31 | 0.56 | 0.63 | 0.92 | 1.63 | 2.11 |
| Phorate ^d | | | | | | | |
| Categorical | 0.97 | 0.28 | 0.46 | 0.55 | 0.93 | 1.46 | 2.15 |
| Trend ^f | 1.00 | 0.31 | 0.57 | 0.64 | 0.92 | 1.62 | 2.15 |
| Terbufos | | | | | | | |
| Categorical | 0.99 | 0.23 | 0.59 | 0.67 | 0.97 | 1.36 | 2.06 |
| Trend ^f | 1.00 | 0.21 | 0.68 | 0.74 | 0.95 | 1.42 | 1.78 |

| Cumulative lifetime days personally mixed or applied ^{b,c} | Mean | Standard deviation | 1st Percentile | 5th Percentile | Median | 95th Percentile | 99th Percentile |
|---|------|--------------------|----------------|----------------|--------|-----------------|-----------------|
| <i>Pyrethroid insecticides</i> | | | | | | | |
| Permethrin (for animals) | | | | | | | |
| Categorical | 0.99 | 0.22 | 0.45 | 0.73 | 0.96 | 1.37 | 1.82 |
| Trend ^f | 1.00 | 0.21 | 0.68 | 0.74 | 0.96 | 1.41 | 1.72 |
| Permethrin (for crops) | | | | | | | |
| Categorical | 1.00 | 0.24 | 0.65 | 0.73 | 0.96 | 1.48 | 1.89 |
| Trend ^f | 1.00 | 0.21 | 0.66 | 0.74 | 0.96 | 1.42 | 1.76 |

Abbreviations: 2,4-D, (2,4-dichlorophenoxy)acetic acid; 2,4,5-T, (2,4,5-trichlorophenoxy)acetic acid; 2,4,5-TP, (RS)-2-(2,4,5-trichlorophenoxy)propionic acid; DDT, 1,1,1-trichloro-2,2-bis(4-chlorophenyl)ethane; EPTC, S-ethyl dipropyl(thiocarbamate); POST, post-enrollment; PRE-B, pre-enrollment both; PRE-E, pre-enrollment enrollment only.

^aWeights adjusted for age at enrollment (modeled with a restricted, quadratic spline with knots at ages 40, 48, and 57 years based on percentiles of the age distribution in cases), ever diagnosed with diabetes, education level, state of residence, not missing covariate data (conditional on age, state, the exposure, and pairwise interaction terms between each covariate and the exposure), and not dropping out of the AHS cohort (conditional on age, diabetes, education, state, the exposure, and pairwise interaction terms between each covariate and the exposure). ^bCategory boundaries set at tertiles of cumulative lifetime days of use for each pesticide among all male private pesticide applicators who used it. ^cFewer than five PRE-E, PRE-B, or POST cases (see footnote a at the bottom of Table S1 for a description of the three case groups) used trichlorfon or ziram at every exposure level. ^dData available only for 11,982 applicators who completed the farmer questionnaire. Weights additionally adjusted for completing the farmer questionnaire (conditional on age, diabetes, education, and state). ^eCategory boundaries set at the median of cumulative lifetime days of use for each pesticide among all male private pesticide applicators who used it. ^fUsed within-category medians. ^gBenomyl is also included in carbamates.

Table S5. Cumulative lifetime days of use of individual pesticides and self-reported depression among male private pesticide applicators in the Agricultural Health Study.

| Cumulative lifetime days personally mixed or applied ^{a,b} | Non-cases [n (%)] | PRE-E ^c Cases [n (%)] | PRE-E ^c IP-weighted OR ^d (95% CI) | PRE-B ^c Cases [n (%)] | PRE-B ^c IP-weighted OR ^d (95% CI) | POST ^c Cases [n (%)] | POST ^c IP-weighted OR ^d (95% CI) | p for difference among ORs ^e |
|---|-------------------|----------------------------------|---|----------------------------------|---|---------------------------------|--|---|
| Total | 19,506 (100) | 474 (100) | | 540 (100) | | 688 (100) | | |
| Fumigants | | | | | | | | |
| Aluminum phosphide ^{f,g} | | | | | | | | |
| 0 (Median = 0.0) | 10,348 (96) | 284 (95) | Reference | 284 (94) | Reference | 345 (94) | Reference | |
| 1-8 (3.5) | 217 (2) | 6 (2) | 0.8 (0.3, 1.9) | 11 (4) | 1.6 (0.8, 3.2) | 13 (4) | 1.6 (0.8, 3.1) | |
| > 8 (14.5) | 168 (2) | 9 (3) | 1.9 (0.9, 3.8) | 6 (2) | 1.0 (0.4, 2.3) | 8 (2) | 1.6 (0.7, 3.5) | 0.48 |
| Missing | 257 | 7 | | 14 | | 5 | | |
| Trend (IQR = 8.8) ^h | | | 1.4 (0.9, 2.2) | | 1.1 (0.7, 1.7) | | 1.4 (0.9, 2.2) | 0.58 |
| Carbon tetrachloride/carbon disulfide (80/20 mix) ^{f,g} | | | | | | | | |
| 0 (Median = 0.0) | 10,210 (95) | 277 (93) | Reference | 284 (95) | Reference | 343 (94) | Reference | |
| 1-12 (3.5) | 378 (4) | 15 (5) | 1.7 (0.9, 3.2) | 12 (4) | 1.3 (0.6, 2.4) | 17 (5) | 1.2 (0.6, 2.2) | |
| > 12 (54.3) | 129 (1) | 7 (2) | 2.1 (1.0, 4.7) | 3 (1) | ⁱ | 6 (2) | 1.2 (0.5, 2.7) | 0.65 |
| Missing | 273 | 7 | | 16 | | 5 | | |
| Trend (IQR = 21.0) ^h | | | 1.4 (1.0, 1.8) | | 1.0 (0.6, 1.6) | | 1.1 (0.8, 1.5) | 0.41 |
| Ethylene dibromide ^f | | | | | | | | |
| 0 (0.0) | 10,294 (96) | 280 (94) | Reference | 285 (94) | Reference | 349 (95) | Reference | |
| 1-8 (3.5) | 151 (1) | 6 (2) | 1.3 (0.5, 3.8) | 6 (2) | 1.7 (0.6, 4.9) | 6 (2) | 1.0 (0.4, 2.8) | |
| 9-28 (15.5) | 174 (2) | 6 (2) | 1.4 (0.6, 3.4) | 4 (1) | ⁱ | 6 (2) | 1.0 (0.4, 2.4) | |
| > 28 (87.5) | 107 (1) | 7 (2) | 2.6 (1.2, 5.8) | 7 (2) | 2.7 (1.2, 6.1) | 5 (1) | 1.7 (0.6, 4.5) | 0.94 |
| Missing | 264 | 7 | | 13 | | 5 | | |
| Trend (46.3) ^h | | | 1.7 (1.1, 2.5) | | 1.7 (1.1, 2.6) | | 1.3 (0.8, 2.2) | 0.69 |
| Methyl bromide | | | | | | | | |
| 0 (0.0) | 16,516 (86) | 392 (84) | Reference | 446 (84) | Reference | 568 (84) | Reference | |
| 1-12 (8.0) | 1,008 (5) | 20 (4) | 1.0 (0.5, 2.1) | 29 (5) | 1.2 (0.6, 2.1) | 43 (6) | 1.5 (1.0, 2.4) | |
| 13-54 (28.0) | 1,221 (6) | 36 (8) | 1.3 (0.9, 1.8) | 35 (7) | 1.1 (0.8, 1.6) | 35 (5) | 0.8 (0.6, 1.2) | |
| > 54 (122.5) | 512 (3) | 18 (4) | 1.6 (1.0, 2.6) | 20 (4) | 1.4 (0.9, 2.2) | 27 (4) | 1.6 (1.1, 2.4) | 0.59 |

| Cumulative lifetime days personally mixed or applied ^{a,b} | Non-cases [n (%)] | PRE-E ^c Cases [n (%)] | PRE-E ^c IP-weighted OR ^d (95% CI) | PRE-B ^c Cases [n (%)] | PRE-B ^c IP-weighted OR ^d (95% CI) | POST ^c Cases [n (%)] | POST ^c IP-weighted OR ^d (95% CI) | p for difference among ORs ^e |
|---|-------------------|----------------------------------|---|----------------------------------|---|---------------------------------|--|---|
| Missing | 249 | 8 | | 10 | | 15 | | |
| Trend (46.3) ^h | | | 1.2 (1.0, 1.4) | | 1.1 (1.0, 1.4) | | 1.2 (1.0, 1.4) | 0.88 |
| Fungicides | | | | | | | | |
| Benomyl ^{f,g,j} | | | | | | | | |
| 0 (0.0) | 9,925 (93) | 272 (92) | Reference | 272 (92) | Reference | 333 (92) | Reference | |
| 1-25 (12.3) | 435 (4) | 15 (5) | 1.3 (0.7, 2.7) | 13 (4) | 1.5 (0.8, 3.1) | 17 (5) | 0.7 (0.4, 1.4) | |
| > 25 (103.3) | 313 (3) | 9 (3) | 1.1 (0.5, 2.2) | 12 (4) | 1.4 (0.8, 2.7) | 14 (4) | 1.5 (0.8, 2.7) | 0.50 |
| Missing | 317 | 10 | | 18 | | 7 | | |
| Trend (52.5) ^h | | | 1.0 (0.7, 1.5) | | 1.2 (0.9, 1.6) | | 1.2 (0.9, 1.7) | 0.72 |
| Captan | | | | | | | | |
| 0 (0.0) | 16,187 (89) | 385 (88) | Reference | 415 (85) | Reference | 552 (89) | Reference | |
| > 0-0.25 (0.3) | 1,194 (7) | 31 (7) | 1.1 (0.7, 1.7) | 50 (10) | 1.6 (1.2, 2.2) | 32 (5) | 0.8 (0.5, 1.1) | |
| > 0.25-8 (3.5) | 187 (1) | 2 (< 1) | ⁱ | 7 (1) | 1.4 (0.6, 3.0) | 9 (1) | 1.5 (0.7, 2.9) | |
| > 8 (64.0) | 590 (3) | 20 (5) | 1.4 (0.9, 2.2) | 19 (4) | 1.3 (0.8, 2.1) | 29 (5) | 1.5 (1.0, 2.3) | 0.06 |
| Missing | 1,348 | 36 | | 49 | | 66 | | |
| Trend (12.5) ^h | | | 1.1 (1.0, 1.2) | | 1.0 (1.0, 1.1) | | 1.1 (1.0, 1.2) | 0.75 |
| Chlorothalonil | | | | | | | | |
| 0 (0.0) | 18,036 (94) | 437 (94) | Reference | 490 (93) | Reference | 622 (92) | Reference | |
| 1-12 (8.0) | 427 (2) | 10 (2) | 0.9 (0.4, 2.0) | 19 (4) | 1.7 (0.9, 3.1) | 19 (3) | 1.2 (0.6, 2.1) | |
| 13-88 (54.3) | 419 (2) | 10 (2) | 1.2 (0.6, 2.4) | 13 (2) | 1.2 (0.7, 2.1) | 14 (2) | 1.2 (0.7, 2.2) | |
| > 88 (200.0) | 394 (2) | 8 (2) | 0.8 (0.4, 1.7) | 6 (1) | 0.6 (0.3, 1.5) | 18 (3) | 1.3 (0.8, 2.1) | 0.67 |
| Missing | 230 | 9 | | 12 | | 15 | | |
| Trend (111.8) ^h | | | 0.9 (0.6, 1.3) | | 0.8 (0.6, 1.2) | | 1.2 (0.9, 1.5) | 0.32 |
| Maneb/mancozeb ^f | | | | | | | | |
| 0 (0.0) | 9,896 (93) | 268 (91) | Reference | 273 (92) | Reference | 330 (90) | Reference | |
| 1-12 (7.0) | 361 (3) | 11 (4) | 1.4 (0.6, 3.1) | 4 (1) | ⁱ | 14 (4) | 1.1 (0.5, 2.3) | |
| 13-56 (50.8) | 238 (2) | 10 (3) | 2.0 (1.0, 3.9) | 7 (2) | 1.0 (0.5, 2.3) | 13 (4) | 1.7 (0.9, 3.0) | |
| > 56 (224.8) | 197 (2) | 5 (2) | 0.9 (0.4, 2.3) | 14 (5) | 3.0 (1.7, 5.3) | 8 (2) | 1.0 (0.5, 2.2) | 0.12 |
| Missing | 298 | 12 | | 17 | | 6 | | |
| Trend (96.3) ^h | | | 1.0 (0.8, 1.4) | | 1.6 (1.2, 2.0) | | 1.1 (0.8, 1.4) | 0.04 |

| Cumulative lifetime days personally mixed or applied ^{a,b} | Non-cases [n (%)] | PRE-E ^c Cases [n (%)] | PRE-E ^c IP-weighted OR ^d (95% CI) | PRE-B ^c Cases [n (%)] | PRE-B ^c IP-weighted OR ^d (95% CI) | POST ^c Cases [n (%)] | POST ^c IP-weighted OR ^d (95% CI) | p for difference among ORs ^e |
|---|-------------------|----------------------------------|---|----------------------------------|---|---------------------------------|--|---|
| Metalaxyf | | | | | | | | |
| 0 (0.0) | 8,790 (83) | 238 (81) | Reference | 242 (81) | Reference | 304 (85) | Reference | |
| 1-12 (3.5) | 1,015 (10) | 32 (11) | 1.0 (0.6, 1.7) | 29 (10) | 1.4 (0.9, 2.2) | 30 (8) | 0.7 (0.5, 1.1) | |
| 13-28 (28.0) | 353 (3) | 10 (3) | 0.9 (0.5, 1.8) | 8 (3) | 0.8 (0.4, 1.8) | 12 (3) | 0.8 (0.4, 1.6) | |
| > 28 (56.0) | 486 (5) | 14 (5) | 1.1 (0.6, 2.0) | 20 (7) | 1.4 (0.9, 2.3) | 10 (3) | 0.7 (0.3, 1.3) | 0.31 |
| Missing | 346 | 12 | | 16 | | 15 | | |
| Trend (50.8) ^h | | | 1.1 (0.7, 1.8) | | 1.3 (0.8, 2.0) | | 0.7 (0.4, 1.2) | 0.20 |
| Herbicides | | | | | | | | |
| Alachlor | | | | | | | | |
| 0 (0.0) | 8,201 (45) | 172 (38) | Reference | 197 (38) | Reference | 272 (42) | Reference | |
| 1-25 (8.8) | 4,619 (25) | 121 (27) | 1.3 (1.0, 1.6) | 137 (27) | 1.2 (0.9, 1.5) | 179 (28) | 1.2 (1.0, 1.5) | |
| 26-109 (56.0) | 3,167 (17) | 94 (21) | 1.5 (1.1, 1.9) | 108 (21) | 1.4 (1.1, 1.8) | 110 (17) | 1.0 (0.8, 1.3) | |
| > 109 (224.8) | 2,440 (13) | 68 (15) | 1.4 (1.0, 1.8) | 72 (14) | 1.2 (0.9, 1.6) | 86 (13) | 1.1 (0.9, 1.4) | 0.42 |
| Missing | 1,079 | 19 | | 26 | | 41 | | |
| Trend (96.0) ^h | | | 1.1 (1.0, 1.2) | | 1.1 (1.0, 1.2) | | 1.0 (0.9, 1.1) | 0.53 |
| Butylate ^f | | | | | | | | |
| 0 (0.0) | 7,616 (71) | 207 (71) | Reference | 200 (66) | Reference | 263 (73) | Reference | |
| 1-9 (8.8) | 1,080 (10) | 32 (11) | 1.2 (0.8, 1.8) | 37 (12) | 1.0 (0.7, 1.5) | 30 (8) | 0.7 (0.4, 1.0) | |
| 10-51 (24.5) | 1,147 (11) | 28 (10) | 0.9 (0.6, 1.4) | 37 (12) | 1.2 (0.8, 1.8) | 45 (12) | 1.1 (0.8, 1.6) | |
| > 51 (108.5) | 846 (8) | 24 (8) | 1.1 (0.7, 1.7) | 28 (9) | 1.2 (0.8, 1.8) | 24 (7) | 0.8 (0.5, 1.3) | 0.27 |
| Missing | 301 | 15 | | 13 | | 9 | | |
| Trend (47.3) ^h | | | 1.0 (0.8, 1.3) | | 1.1 (0.9, 1.3) | | 0.9 (0.8, 1.1) | 0.46 |
| Chlorimuron-ethyl ^f | | | | | | | | |
| 0 (0.0) | 7,240 (68) | 210 (71) | Reference | 185 (61) | Reference | 249 (68) | Reference | |
| 1-9 (8.8) | 2,264 (21) | 51 (17) | 0.7 (0.5, 1.0) | 82 (27) | 1.4 (1.1, 1.9) | 76 (21) | 1.0 (0.7, 1.3) | |
| 10-25 (24.5) | 673 (6) | 16 (5) | 0.8 (0.5, 1.4) | 21 (7) | 1.2 (0.8, 2.0) | 26 (7) | 1.2 (0.7, 1.8) | |
| > 25 (56.0) | 530 (5) | 20 (7) | 1.3 (0.8, 2.1) | 13 (4) | 0.9 (0.5, 1.6) | 14 (4) | 0.7 (0.4, 1.3) | 0.03 |
| Missing | 283 | 9 | | 14 | | 6 | | |
| Trend (15.8) ^h | | | 1.0 (0.9, 1.2) | | 1.0 (0.9, 1.1) | | 0.9 (0.8, 1.1) | 0.64 |

| Cumulative lifetime days personally mixed or applied ^{a,b} | Non-cases [n (%)] | PRE-E ^c Cases [n (%)] | PRE-E ^c IP-weighted OR ^d (95% CI) | PRE-B ^c Cases [n (%)] | PRE-B ^c IP-weighted OR ^d (95% CI) | POST ^c Cases [n (%)] | POST ^c IP-weighted OR ^d (95% CI) | p for difference among ORs ^e |
|---|-------------------|----------------------------------|---|----------------------------------|---|---------------------------------|--|---|
| Dicamba | | | | | | | | |
| 0 (0.0) | 8,367 (45) | 209 (46) | Reference | 217 (43) | Reference | 280 (44) | Reference | |
| 1-20 (8.8) | 4,315 (23) | 93 (20) | 0.8 (0.6, 1.1) | 134 (27) | 1.0 (0.8, 1.3) | 146 (23) | 0.9 (0.7, 1.2) | |
| 21-56 (38.8) | 3,010 (16) | 74 (16) | 1.0 (0.7, 1.3) | 77 (15) | 1.0 (0.8, 1.3) | 100 (16) | 1.0 (0.8, 1.3) | |
| > 56 (116.0) | 2,715 (15) | 78 (17) | 1.1 (0.9, 1.5) | 77 (15) | 1.1 (0.9, 1.5) | 107 (17) | 1.2 (1.0, 1.5) | 0.94 |
| Missing | 1,099 | 20 | | 35 | | 55 | | |
| Trend (99.8) ^h | | | 1.1 (0.9, 1.4) | | 1.1 (0.9, 1.3) | | 1.1 (0.9, 1.4) | 0.93 |
| EPTC | | | | | | | | |
| 0 (0.0) | 14,440 (79) | 335 (75) | Reference | 397 (80) | Reference | 492 (77) | Reference | |
| 1-9 (8.8) | 1,800 (10) | 50 (11) | 1.2 (0.9, 1.7) | 47 (9) | 1.0 (0.7, 1.4) | 77 (12) | 1.2 (0.9, 1.6) | |
| 10-25 (24.5) | 920 (5) | 24 (5) | 1.1 (0.7, 1.7) | 26 (5) | 1.1 (0.7, 1.7) | 38 (6) | 1.2 (0.8, 1.7) | |
| > 25 (103.3) | 1,212 (7) | 38 (9) | 1.4 (1.0, 1.9) | 28 (6) | 0.9 (0.6, 1.3) | 36 (6) | 0.9 (0.6, 1.2) | 0.55 |
| Missing | 1,134 | 27 | | 42 | | 45 | | |
| Trend (47.3) ^h | | | 1.1 (1.0, 1.3) | | 1.0 (0.8, 1.1) | | 0.9 (0.8, 1.1) | 0.15 |
| Glyphosate | | | | | | | | |
| 0 (0.0) | 4,371 (23) | 94 (20) | Reference | 113 (21) | Reference | 143 (21) | Reference | |
| 1-20 (8.8) | 6,219 (32) | 135 (29) | 1.1 (0.8, 1.4) | 203 (38) | 1.3 (1.0, 1.6) | 219 (32) | 1.0 (0.8, 1.3) | |
| 21-56 (38.8) | 4,497 (23) | 116 (25) | 1.2 (0.9, 1.6) | 107 (20) | 0.9 (0.6, 1.1) | 150 (22) | 1.1 (0.8, 1.4) | |
| > 56 (116.0) | 4,183 (22) | 124 (26) | 1.4 (1.0, 1.8) | 112 (21) | 1.0 (0.8, 1.3) | 163 (24) | 1.2 (0.9, 1.5) | 0.04 |
| Missing | 236 | 5 | | 5 | | 13 | | |
| Trend (99.8) ^h | | | 1.3 (1.1, 1.6) | | 0.9 (0.7, 1.1) | | 1.1 (1.0, 1.4) | 0.02 |
| Imazethapyr | | | | | | | | |
| 0 (0.0) | 10,075 (55) | 240 (54) | Reference | 288 (58) | Reference | 347 (54) | Reference | |
| 1-9 (8.8) | 3,733 (20) | 91 (20) | 1.1 (0.8, 1.5) | 97 (20) | 1.0 (0.7, 1.4) | 129 (20) | 1.2 (0.9, 1.6) | |
| 10-25 (24.5) | 2,639 (14) | 61 (14) | 1.0 (0.7, 1.3) | 72 (14) | 1.0 (0.7, 1.3) | 105 (16) | 1.2 (0.9, 1.5) | |
| > 25 (56.0) | 1,975 (11) | 53 (12) | 1.1 (0.8, 1.5) | 40 (8) | 0.7 (0.5, 1.0) | 66 (10) | 1.0 (0.7, 1.3) | 0.57 |
| Missing | 1,084 | 29 | | 43 | | 41 | | |
| Trend (30.0) ^h | | | 1.0 (0.9, 1.2) | | 0.9 (0.7, 1.0) | | 1.0 (0.9, 1.1) | 0.28 |

| Cumulative lifetime days personally mixed or applied ^{a,b} | Non-cases [n (%)] | PRE-E ^c Cases [n (%)] | PRE-E ^c IP-weighted OR ^d (95% CI) | PRE-B ^c Cases [n (%)] | PRE-B ^c IP-weighted OR ^d (95% CI) | POST ^c Cases [n (%)] | POST ^c IP-weighted OR ^d (95% CI) | p for difference among ORs ^e |
|---|-------------------|----------------------------------|---|----------------------------------|---|---------------------------------|--|---|
| Metolachlor | | | | | | | | |
| 0 (0.0) | 9,560 (52) | 224 (50) | Reference | 281 (56) | Reference | 341 (52) | Reference | |
| 1-25 (8.8) | 4,309 (23) | 101 (22) | 1.1 (0.8, 1.4) | 116 (23) | 0.9 (0.7, 1.2) | 139 (21) | 1.0 (0.8, 1.2) | |
| 26-56 (56.0) | 1,898 (10) | 47 (10) | 1.0 (0.8, 1.4) | 50 (10) | 0.9 (0.7, 1.2) | 78 (12) | 1.1 (0.9, 1.5) | |
| > 56 (116.0) | 2,732 (15) | 79 (18) | 1.2 (0.9, 1.6) | 55 (11) | 0.7 (0.5, 1.0) | 92 (14) | 1.0 (0.8, 1.2) | 0.26 |
| Missing | 977 | 23 | | 38 | | 38 | | |
| Trend (88.5) ^h | | | 1.1 (0.9, 1.4) | | 0.8 (0.6, 1.0) | | 1.0 (0.8, 1.2) | 0.03 |
| Paraquat ^f | | | | | | | | |
| 0 (0.0) | 9,054 (84) | 245 (82) | Reference | 253 (85) | Reference | 309 (86) | Reference | |
| 1-9 (8.8) | 911 (9) | 30 (10) | 1.6 (1.0, 2.4) | 32 (11) | 1.2 (0.7, 1.8) | 25 (7) | 0.7 (0.4, 1.2) | |
| 10-25 (24.5) | 309 (3) | 11 (4) | 1.2 (0.6, 2.4) | 3 (1) | | 12 (3) | 0.9 (0.5, 1.6) | |
| > 25 (108.5) | 442 (4) | 11 (4) | 0.9 (0.5, 1.8) | 10 (3) | 0.7 (0.3, 1.3) | 15 (4) | 1.1 (0.6, 1.9) | 0.13 |
| Missing | 274 | 9 | | 17 | | 10 | | |
| Trend (42.0) ^h | | | 1.0 (0.8, 1.3) | | 0.8 (0.6, 1.1) | | 1.0 (0.8, 1.3) | 0.56 |
| Pendimethalin ^f | | | | | | | | |
| 0 (0.0) | 6,752 (63) | 182 (61) | Reference | 192 (64) | Reference | 249 (68) | Reference | |
| 1-9 (8.8) | 1,804 (17) | 48 (16) | 1.0 (0.7, 1.4) | 63 (21) | 1.2 (0.9, 1.6) | 55 (15) | 0.7 (0.6, 1.0) | |
| 10-25 (24.5) | 1,000 (9) | 28 (9) | 1.0 (0.6, 1.5) | 18 (6) | 0.5 (0.3, 0.9) | 30 (8) | 0.8 (0.5, 1.2) | |
| > 25 (56.0) | 1,138 (11) | 38 (13) | 1.2 (0.8, 1.8) | 26 (9) | 0.8 (0.5, 1.2) | 30 (8) | 0.7 (0.5, 1.1) | 0.06 |
| Missing | 296 | 10 | | 16 | | 7 | | |
| Trend (42.0) ^h | | | 1.1 (0.9, 1.5) | | 0.8 (0.5, 1.1) | | 0.8 (0.6, 1.0) | 0.09 |
| Petroleum oil ^f | | | | | | | | |
| 0 (0.0) | 8,295 (78) | 223 (76) | Reference | 218 (74) | Reference | 272 (75) | Reference | |
| 1-20 (8.8) | 1,002 (9) | 22 (8) | 0.9 (0.6, 1.5) | 34 (11) | 1.2 (0.8, 1.7) | 34 (9) | 1.1 (0.7, 1.6) | |
| 21-56 (38.8) | 653 (6) | 16 (5) | 0.8 (0.5, 1.5) | 18 (6) | 1.0 (0.6, 1.6) | 31 (9) | 1.4 (1.0, 2.1) | |
| > 56 (224.8) | 680 (6) | 32 (11) | 1.6 (1.1, 2.4) | 26 (9) | 1.4 (0.9, 2.1) | 25 (7) | 1.1 (0.7, 1.7) | 0.50 |
| Missing | 360 | 13 | | 19 | | 9 | | |
| Trend (99.8) ^h | | | 1.2 (1.0, 1.5) | | 1.1 (0.9, 1.4) | | 1.1 (0.9, 1.3) | 0.51 |

| Cumulative lifetime days personally mixed or applied ^{a,b} | Non-cases [n (%)] | PRE-E ^c Cases [n (%)] | PRE-E ^c IP-weighted OR ^d (95% CI) | PRE-B ^c Cases [n (%)] | PRE-B ^c IP-weighted OR ^d (95% CI) | POST ^c Cases [n (%)] | POST ^c IP-weighted OR ^d (95% CI) | p for difference among ORs ^e |
|---|-------------------|----------------------------------|---|----------------------------------|---|---------------------------------|--|---|
| Trifluralin | | | | | | | | |
| 0 (0.0) | 8,343 (45) | 183 (41) | Reference | 218 (43) | Reference | 285 (45) | Reference | |
| 1-25 (20.0) | 3,512 (19) | 84 (19) | 1.2 (0.9, 1.6) | 109 (21) | 1.2 (0.9, 1.5) | 111 (17) | 1.1 (0.8, 1.4) | |
| 26-109 (56.0) | 3,953 (21) | 94 (21) | 1.1 (0.9, 1.4) | 111 (22) | 1.1 (0.9, 1.4) | 147 (23) | 1.1 (0.9, 1.3) | |
| > 109 (224.8) | 2,600 (14) | 86 (19) | 1.5 (1.1, 1.9) | 73 (14) | 1.1 (0.8, 1.4) | 94 (15) | 1.1 (0.8, 1.4) | 0.61 |
| Missing | 1,098 | 27 | | 29 | | 51 | | |
| Trend (91.5) ^h | | | 1.2 (1.0, 1.3) | | 1.0 (0.9, 1.1) | | 1.0 (0.9, 1.1) | 0.19 |
| Phenoxy herbicides | | | | | | | | |
| 2,4-D | | | | | | | | |
| 0 (0.0) | 3,976 (21) | 91 (20) | Reference | 95 (18) | Reference | 151 (23) | Reference | |
| 1-39 (20.0) | 5,950 (31) | 100 (22) | 0.7 (0.5, 1.0) | 174 (33) | 1.2 (0.9, 1.6) | 210 (31) | 1.0 (0.8, 1.2) | |
| 40-116 (87.5) | 4,629 (24) | 135 (29) | 1.3 (1.0, 1.7) | 133 (25) | 1.3 (1.0, 1.7) | 161 (24) | 0.9 (0.7, 1.2) | |
| > 116 (245.0) | 4,575 (24) | 137 (30) | 1.3 (1.0, 1.7) | 131 (25) | 1.3 (1.0, 1.6) | 147 (22) | 0.9 (0.7, 1.1) | < 0.01 |
| Missing | 376 | 11 | | 7 | | 19 | | |
| Trend (154.0) ^h | | | 1.3 (1.1, 1.5) | | 1.1 (0.9, 1.2) | | 0.9 (0.8, 1.1) | < 0.01 |
| 2,4,5-T ^f | | | | | | | | |
| 0 (0.0) | 8,541 (80) | 204 (70) | Reference | 223 (74) | Reference | 290 (81) | Reference | |
| 1-9 (8.8) | 1,240 (12) | 52 (18) | 1.9 (1.3, 2.7) | 42 (14) | 1.4 (0.9, 2.1) | 44 (12) | 1.1 (0.7, 1.5) | |
| 10-25 (22.3) | 382 (4) | 21 (7) | 2.2 (1.4, 3.6) | 15 (5) | 1.5 (0.9, 2.6) | 11 (3) | 0.8 (0.4, 1.5) | |
| > 25 (62.5) | 518 (5) | 16 (5) | 1.3 (0.8, 2.3) | 20 (7) | 1.5 (0.9, 2.4) | 14 (4) | 0.7 (0.4, 1.3) | 0.03 |
| Missing | 309 | 13 | | 15 | | 12 | | |
| Trend (30.0) ^h | | | 1.3 (1.1, 1.6) | | 1.3 (1.0, 1.6) | | 0.9 (0.7, 1.1) | 0.03 |
| 2,4,5-TP ^{f,g} | | | | | | | | |
| 0 (0.0) | 10,114 (95) | 268 (91) | Reference | 281 (94) | Reference | 338 (94) | Reference | |
| 1-20 (8.8) | 319 (3) | 13 (4) | 1.9 (1.0, 3.5) | 12 (4) | 1.5 (0.8, 3.1) | 17 (5) | 1.7 (1.0, 3.0) | |
| > 20 (56.0) | 223 (2) | 12 (4) | 2.1 (1.1, 3.9) | 7 (2) | 1.1 (0.5, 2.5) | 5 (1) | 0.6 (0.2, 1.5) | 0.23 |
| Missing | 334 | 13 | | 15 | | 11 | | |
| Trend (47.4) ^h | | | 2.0 (1.2, 3.2) | | 1.2 (0.6, 2.2) | | 0.8 (0.4, 1.5) | 0.05 |

| Cumulative lifetime days personally mixed or applied ^{a,b} | Non-cases [n (%)] | PRE-E ^c Cases [n (%)] | PRE-E ^c IP-weighted OR ^d (95% CI) | PRE-B ^c Cases [n (%)] | PRE-B ^c IP-weighted OR ^d (95% CI) | POST ^c Cases [n (%)] | POST ^c IP-weighted OR ^d (95% CI) | p for difference among ORs ^e |
|---|-------------------|----------------------------------|---|----------------------------------|---|---------------------------------|--|---|
| <i>Triazine herbicides</i> | | | | | | | | |
| Atrazine | | | | | | | | |
| 0 (0.0) | 4,854 (25) | 100 (21) | Reference | 122 (23) | Reference | 173 (26) | Reference | |
| 1-25 (8.8) | 4,748 (25) | 105 (22) | 1.0 (0.8, 1.4) | 120 (23) | 1.0 (0.7, 1.2) | 179 (26) | 1.1 (0.9, 1.4) | |
| 26-109 (56.0) | 5,353 (28) | 131 (28) | 1.2 (0.9, 1.6) | 175 (33) | 1.3 (1.0, 1.7) | 159 (23) | 0.8 (0.7, 1.1) | |
| > 109 (224.8) | 4,285 (22) | 133 (28) | 1.5 (1.2, 2.0) | 116 (22) | 1.0 (0.8, 1.4) | 166 (25) | 1.1 (0.9, 1.4) | < 0.01 |
| Missing | 266 | 5 | | 7 | | 11 | | |
| Trend (158.5) ^h | | | 1.3 (1.1, 1.5) | | 1.0 (0.9, 1.2) | | 1.1 (0.9, 1.2) | 0.11 |
| Cyanazine | | | | | | | | |
| 0 (0.0) | 10,284 (55) | 222 (49) | Reference | 259 (50) | Reference | 350 (54) | Reference | |
| 1-20 (8.8) | 3,375 (18) | 77 (17) | 1.0 (0.7, 1.4) | 92 (18) | 1.0 (0.7, 1.3) | 128 (20) | 1.2 (1.0, 1.6) | |
| 21-56 (38.8) | 2,611 (14) | 79 (17) | 1.4 (1.1, 1.9) | 89 (17) | 1.3 (1.0, 1.7) | 90 (14) | 1.0 (0.8, 1.3) | |
| > 56 (116.0) | 2,276 (12) | 74 (16) | 1.5 (1.2, 2.0) | 75 (15) | 1.3 (1.0, 1.8) | 82 (13) | 1.1 (0.8, 1.3) | 0.05 |
| Missing | 960 | 22 | | 25 | | 38 | | |
| Trend (99.8) ^h | | | 1.4 (1.2, 1.8) | | 1.3 (1.0, 1.6) | | 1.0 (0.8, 1.3) | 0.10 |
| Metribuzin ^f | | | | | | | | |
| 0 (0.0) | 6,347 (59) | 169 (57) | Reference | 160 (54) | Reference | 226 (62) | Reference | |
| 1-9 (8.8) | 2,099 (20) | 65 (22) | 1.3 (0.9, 1.9) | 67 (23) | 1.0 (0.7, 1.4) | 57 (16) | 0.8 (0.5, 1.1) | |
| 10-25 (24.5) | 1,110 (10) | 27 (9) | 0.9 (0.6, 1.3) | 35 (12) | 1.3 (0.9, 1.8) | 44 (12) | 1.1 (0.8, 1.5) | |
| > 25 (56.0) | 1,121 (11) | 35 (12) | 1.2 (0.8, 1.8) | 34 (11) | 1.2 (0.8, 1.8) | 35 (10) | 0.9 (0.6, 1.3) | 0.23 |
| Missing | 313 | 10 | | 19 | | 9 | | |
| Trend (30.0) ^h | | | 1.1 (0.9, 1.3) | | 1.1 (0.9, 1.3) | | 0.9 (0.8, 1.1) | 0.47 |
| <i>Insecticides</i> | | | | | | | | |
| Carbamates ^j | | | | | | | | |
| Aldicarb ^{f,g} | | | | | | | | |
| 0 (0.0) | 9,974 (93) | 272 (92) | Reference | 280 (94) | Reference | 332 (91) | Reference | |
| 1-25 (8.8) | 425 (4) | 17 (6) | 1.8 (0.8, 3.7) | 15 (5) | 1.9 (0.9, 4.1) | 22 (6) | 2.2 (1.2, 4.1) | |
| > 25 (108.5) | 298 (3) | 6 (2) | 1.1 (0.5, 2.7) | 4 (1) | ⁱ | 9 (2) | 1.1 (0.5, 2.2) | 0.66 |
| Missing | 293 | 11 | | 16 | | 8 | | |
| Trend (99.8) ^h | | | 1.1 (0.5, 2.4) | | 0.5 (0.2, 1.2) | | 1.1 (0.6, 2.0) | 0.32 |

| Cumulative lifetime days personally mixed or applied ^{a,b} | Non-cases [n (%)] | PRE-E ^c Cases [n (%)] | PRE-E ^c IP-weighted OR ^d (95% CI) | PRE-B ^c Cases [n (%)] | PRE-B ^c IP-weighted OR ^d (95% CI) | POST ^c Cases [n (%)] | POST ^c IP-weighted OR ^d (95% CI) | p for difference among ORs ^e |
|---|-------------------|----------------------------------|---|----------------------------------|---|---------------------------------|--|---|
| Carbaryl ^f | | | | | | | | |
| 0 (0.0) | 6,033 (57) | 160 (54) | Reference | 154 (52) | Reference | 206 (57) | Reference | |
| 1-9 (8.8) | 1,892 (18) | 58 (20) | 1.2 (0.9, 1.7) | 68 (23) | 1.4 (1.0, 1.9) | 62 (17) | 0.9 (0.7, 1.3) | |
| 10-56 (24.5) | 1,521 (14) | 32 (11) | 1.0 (0.7, 1.6) | 32 (11) | 0.8 (0.5, 1.3) | 50 (14) | 0.9 (0.7, 1.3) | |
| > 56 (175.0) | 1,190 (11) | 46 (16) | 1.7 (1.2, 2.4) | 41 (14) | 1.4 (1.0, 2.0) | 44 (12) | 1.1 (0.7, 1.5) | 0.36 |
| Missing | 354 | 10 | | 20 | | 9 | | |
| Trend (99.8) ^h | | | 1.3 (1.1, 1.6) | | 1.2 (1.0, 1.5) | | 1.1 (0.9, 1.3) | 0.29 |
| Carbofuran | | | | | | | | |
| 0 (0.0) | 12,998 (71) | 296 (67) | Reference | 331 (65) | Reference | 468 (73) | Reference | |
| 1-9 (8.8) | 2,245 (12) | 53 (12) | 1.0 (0.8, 1.4) | 77 (15) | 1.3 (1.0, 1.7) | 69 (11) | 0.9 (0.7, 1.2) | |
| 10-51 (24.5) | 1,970 (11) | 57 (13) | 1.3 (1.0, 1.7) | 63 (12) | 1.3 (1.0, 1.7) | 71 (11) | 1.0 (0.8, 1.3) | |
| > 51 (116.0) | 1,224 (7) | 39 (9) | 1.3 (0.9, 1.9) | 35 (7) | 1.1 (0.8, 1.6) | 36 (6) | 0.9 (0.6, 1.2) | 0.23 |
| Missing | 1,069 | 29 | | 34 | | 44 | | |
| Trend (47.3) ^h | | | 1.1 (1.0, 1.3) | | 1.0 (0.9, 1.2) | | 0.9 (0.8, 1.1) | 0.14 |
| Organochlorine insecticides | | | | | | | | |
| Aldrin ^f | | | | | | | | |
| 0 (0.0) | 8,722 (82) | 214 (73) | Reference | 218 (74) | Reference | 293 (81) | Reference | |
| 1-9 (8.8) | 904 (8) | 31 (11) | 1.7 (1.0, 2.8) | 31 (10) | 1.5 (0.9, 2.5) | 44 (12) | 1.6 (1.1, 2.4) | |
| 10-25 (24.5) | 576 (5) | 35 (12) | 2.5 (1.7, 3.7) | 27 (9) | 1.9 (1.3, 3.0) | 15 (4) | 0.7 (0.4, 1.2) | |
| > 25 (56.0) | 464 (4) | 15 (5) | 1.2 (0.7, 2.1) | 20 (7) | 1.7 (1.1, 2.7) | 9 (2) | 0.6 (0.3, 1.1) | < 0.01 |
| Missing | 324 | 11 | | 19 | | 10 | | |
| Trend (15.8) ^h | | | 1.2 (1.1, 1.3) | | 1.2 (1.1, 1.4) | | 0.9 (0.8, 1.0) | < 0.01 |
| Chlordane ^f | | | | | | | | |
| 0 (0.0) | 8,480 (80) | 205 (70) | Reference | 224 (76) | Reference | 285 (79) | Reference | |
| 1-9 (8.8) | 1,464 (14) | 58 (20) | 2.0 (1.4, 2.8) | 45 (15) | 1.1 (0.7, 1.6) | 53 (15) | 1.3 (0.9, 1.8) | |
| 10-20 (20.0) | 189 (2) | 4 (1) | ⁱ | 6 (2) | 1.5 (0.6, 3.4) | 8 (2) | 1.1 (0.5, 2.4) | |
| > 20 (50.8) | 522 (5) | 27 (9) | 2.0 (1.3, 3.0) | 21 (7) | 1.7 (1.0, 2.7) | 15 (4) | 0.7 (0.4, 1.3) | 0.01 |
| Missing | 335 | 12 | | 19 | | 10 | | |
| Trend (22.0) ^h | | | 1.4 (1.2, 1.6) | | 1.3 (1.0, 1.6) | | 0.9 (0.7, 1.1) | 0.01 |

| Cumulative lifetime days personally mixed or applied ^{a,b} | Non-cases [n (%)] | PRE-E ^c Cases [n (%)] | PRE-E ^c IP-weighted OR ^d (95% CI) | PRE-B ^c Cases [n (%)] | PRE-B ^c IP-weighted OR ^d (95% CI) | POST ^c Cases [n (%)] | POST ^c IP-weighted OR ^d (95% CI) | p for difference among ORs ^e |
|---|-------------------|----------------------------------|---|----------------------------------|---|---------------------------------|--|---|
| DDT ^f | | | | | | | | |
| 0 (Median = 0.0) | 8,159 (77) | 205 (70) | Reference | 219 (74) | Reference | 284 (80) | Reference | |
| 1-9 (8.8) | 1,147 (11) | 34 (12) | 1.1 (0.7, 1.7) | 29 (10) | 1.3 (0.8, 2.3) | 34 (10) | 1.1 (0.7, 1.9) | |
| 10-39 (24.5) | 571 (5) | 21 (7) | 1.6 (1.0, 2.5) | 17 (6) | 1.1 (0.7, 1.8) | 17 (5) | 0.9 (0.6, 1.6) | |
| > 39 (108.5) | 761 (7) | 31 (11) | 1.7 (1.1, 2.5) | 30 (10) | 1.7 (1.1, 2.5) | 23 (6) | 0.8 (0.5, 1.3) | 0.19 |
| Missing | 352 | 15 | | 20 | | 13 | | |
| Trend (47.3) ^h | | | 1.3 (1.1, 1.5) | | 1.2 (1.0, 1.5) | | 0.9 (0.8, 1.1) | 0.02 |
| Dieldrin ^{f,g} | | | | | | | | |
| 0 (0.0) | 10,219 (96) | 276 (95) | Reference | 284 (96) | Reference | 350 (96) | Reference | |
| 1-9 (8.8) | 279 (3) | 9 (3) | 1.0 (0.5, 2.0) | 10 (3) | 1.2 (0.5, 3.0) | 9 (2) | 1.3 (0.5, 3.1) | |
| > 9 (24.5) | 152 (1) | 7 (2) | 1.5 (0.7, 3.3) | 3 (1) | | 5 (1) | 1.3 (0.5, 3.4) | 0.70 |
| Missing | 340 | 14 | | 18 | | 7 | | |
| Trend (15.8) ^h | | | 1.4 (0.9, 2.1) | | 0.9 (0.5, 1.5) | | 1.1 (0.6, 2.0) | 0.50 |
| Heptachlor ^f | | | | | | | | |
| 0 (0.0) | 9,283 (87) | 226 (76) | Reference | 253 (84) | Reference | 314 (86) | Reference | |
| 1-9 (8.8) | 731 (7) | 28 (9) | 1.6 (0.9, 2.6) | 27 (9) | 1.4 (0.8, 2.5) | 25 (7) | 1.3 (0.7, 2.1) | |
| 10-25 (24.5) | 387 (4) | 23 (8) | 2.5 (1.6, 3.9) | 14 (5) | 1.3 (0.8, 2.3) | 14 (4) | 1.1 (0.6, 1.8) | |
| > 25 (56.0) | 291 (3) | 21 (7) | 3.1 (1.9, 5.0) | 7 (2) | 0.8 (0.4, 1.8) | 11 (3) | 1.2 (0.6, 2.2) | 0.01 |
| Missing | 298 | 8 | | 14 | | 7 | | |
| Trend (15.8) ^h | | | 1.4 (1.3, 1.6) | | 1.0 (0.9, 1.2) | | 1.1 (0.9, 1.2) | < 0.01 |
| Lindane ^f | | | | | | | | |
| 0 (0.0) | 9,029 (85) | 232 (79) | Reference | 223 (76) | Reference | 287 (80) | Reference | |
| 1-9 (8.8) | 740 (7) | 19 (6) | 1.0 (0.6, 1.7) | 40 (14) | 2.0 (1.4, 3.0) | 34 (9) | 1.4 (0.9, 2.2) | |
| 10-25 (20.0) | 340 (3) | 20 (7) | 2.1 (1.3, 3.4) | 7 (2) | 0.8 (0.3, 1.7) | 16 (4) | 1.6 (0.9, 2.7) | |
| > 25 (103.3) | 527 (5) | 24 (8) | 1.7 (1.1, 2.6) | 25 (8) | 2.0 (1.3, 3.1) | 23 (6) | 1.4 (0.9, 2.1) | 0.07 |
| Missing | 354 | 11 | | 20 | | 11 | | |
| Trend (47.3) ^h | | | 1.3 (1.1, 1.6) | | 1.4 (1.1, 1.7) | | 1.2 (1.0, 1.4) | 0.54 |
| Toxaphene ^f | | | | | | | | |
| 0 (0.0) | 9,422 (88) | 247 (83) | Reference | 255 (85) | Reference | 318 (87) | Reference | |
| 1-9 (8.8) | 655 (6) | 26 (9) | 1.4 (0.9, 2.2) | 22 (7) | 1.2 (0.8, 2.0) | 21 (6) | 0.9 (0.6, 1.6) | |

| Cumulative lifetime days personally mixed or applied ^{a,b} | Non-cases [n (%)] | PRE-E ^c Cases [n (%)] | PRE-E ^c IP-weighted OR ^d (95% CI) | PRE-B ^c Cases [n (%)] | PRE-B ^c IP-weighted OR ^d (95% CI) | POST ^c Cases [n (%)] | POST ^c IP-weighted OR ^d (95% CI) | p for difference among ORs ^e |
|---|-------------------|----------------------------------|---|----------------------------------|---|---------------------------------|--|---|
| 10-25 (24.5) | 283 (3) | 12 (4) | 1.4 (0.7, 2.5) | 11 (4) | 1.5 (0.8, 2.9) | 15 (4) | 1.5 (0.9, 2.7) | |
| > 25 (108.5) | 331 (3) | 14 (5) | 1.5 (0.9, 2.7) | 12 (4) | 1.5 (0.8, 2.8) | 11 (3) | 0.9 (0.5, 1.7) | 0.76 |
| Missing | 299 | 7 | | 15 | | 6 | | |
| Trend (42.0) ^h | | | 1.2 (1.0, 1.5) | | 1.2 (0.9, 1.5) | | 1.0 (0.8, 1.2) | 0.32 |
| <i>Organophosphate insecticides</i> | | | | | | | | |
| Chlorpyrifos | | | | | | | | |
| 0 (0.0) | 10,913 (57) | 252 (54) | Reference | 267 (50) | Reference | 379 (56) | Reference | |
| 1-20 (8.8) | 3,707 (19) | 107 (23) | 1.3 (1.0, 1.7) | 118 (22) | 1.4 (1.1, 1.7) | 132 (20) | 1.0 (0.8, 1.3) | |
| 21-56 (38.8) | 2,850 (15) | 62 (13) | 0.9 (0.7, 1.2) | 94 (18) | 1.3 (1.0, 1.7) | 107 (16) | 1.1 (0.9, 1.4) | |
| > 56 (116.0) | 1,785 (9) | 47 (10) | 1.1 (0.8, 1.6) | 54 (10) | 1.1 (0.8, 1.5) | 57 (8) | 0.9 (0.7, 1.2) | 0.24 |
| Missing | 251 | 6 | | 7 | | 13 | | |
| Trend (47.3) ^h | | | 1.0 (0.9, 1.2) | | 1.1 (0.9, 1.2) | | 1.0 (0.9, 1.1) | 0.67 |
| Coumaphos | | | | | | | | |
| 0 (0.0) | 16,483 (90) | 387 (88) | Reference | 437 (88) | Reference | 576 (92) | Reference | |
| 1-9 (8.8) | 777 (4) | 16 (4) | 0.8 (0.5, 1.4) | 26 (5) | 1.2 (0.8, 1.8) | 27 (4) | 0.9 (0.6, 1.3) | |
| 10-39 (24.5) | 490 (3) | 18 (4) | 1.5 (1.0, 2.5) | 18 (4) | 1.4 (0.9, 2.3) | 16 (3) | 0.9 (0.5, 1.5) | |
| > 39 (116.0) | 480 (3) | 18 (4) | 1.5 (0.9, 2.4) | 16 (3) | 1.2 (0.7, 2.1) | 8 (1) | 0.5 (0.2, 1.0) | 0.10 |
| Missing | 1,276 | 35 | | 43 | | 61 | | |
| Trend (47.3) ^h | | | 1.2 (1.0, 1.4) | | 1.1 (0.9, 1.4) | | 0.8 (0.6, 1.0) | 0.02 |
| Diazinon ^f | | | | | | | | |
| 0 (0.0) | 8,386 (79) | 229 (78) | Reference | 209 (71) | Reference | 271 (75) | Reference | |
| 1-9 (8.8) | 1,107 (10) | 34 (12) | 1.2 (0.8, 1.8) | 44 (15) | 1.7 (1.2, 2.5) | 43 (12) | 1.2 (0.8, 1.7) | |
| 10-25 (24.5) | 536 (5) | 13 (4) | 0.7 (0.4, 1.3) | 14 (5) | 1.0 (0.5, 1.8) | 18 (5) | 1.2 (0.7, 1.9) | |
| > 25 (103.3) | 600 (6) | 19 (6) | 1.2 (0.7, 1.9) | 27 (9) | 1.9 (1.2, 2.9) | 29 (8) | 1.5 (1.0, 2.2) | 0.42 |
| Missing | 361 | 11 | | 21 | | 10 | | |
| Trend (42.0) ^h | | | 1.1 (0.9, 1.3) | | 1.3 (1.1, 1.5) | | 1.2 (1.0, 1.4) | 0.39 |
| Dichlorvos | | | | | | | | |
| 0 (0.0) | 16,229 (88) | 386 (86) | Reference | 410 (82) | Reference | 549 (85) | Reference | |
| 1-20 (8.8) | 847 (5) | 22 (5) | 1.3 (0.8, 2.1) | 30 (6) | 1.3 (0.8, 1.9) | 36 (6) | 1.3 (0.9, 1.9) | |

| Cumulative lifetime days personally mixed or applied ^{a,b} | Non-cases [n (%)] | PRE-E ^c Cases [n (%)] | PRE-E ^c IP-weighted OR ^d (95% CI) | PRE-B ^c Cases [n (%)] | PRE-B ^c IP-weighted OR ^d (95% CI) | POST ^c Cases [n (%)] | POST ^c IP-weighted OR ^d (95% CI) | p for difference among ORs ^e |
|---|-------------------|----------------------------------|---|----------------------------------|---|---------------------------------|--|---|
| 21-116 (56.0) | 759 (4) | 24 (5) | 1.3 (0.9, 2.0) | 37 (7) | 2.0 (1.4, 2.9) | 32 (5) | 1.3 (0.9, 1.8) | |
| > 116 (457.3) | 634 (3) | 15 (3) | 0.9 (0.6, 1.6) | 24 (5) | 1.5 (1.0, 2.2) | 28 (4) | 1.3 (0.9, 1.9) | 0.45 |
| Missing | 1,037 | 27 | | 39 | | 43 | | |
| Trend (216.0) ^h | | | 1.0 (0.8, 1.2) | | 1.2 (1.0, 1.5) | | 1.1 (0.9, 1.4) | 0.35 |
| Fonofos | | | | | | | | |
| 0 (0.0) | 14,176 (77) | 319 (71) | Reference | 369 (72) | Reference | 498 (78) | Reference | |
| 1-20 (8.8) | 1,791 (10) | 50 (11) | 1.2 (0.9, 1.7) | 57 (11) | 1.2 (0.9, 1.6) | 54 (8) | 0.9 (0.6, 1.2) | |
| 21-51 (24.5) | 1,206 (7) | 45 (10) | 1.6 (1.1, 2.2) | 39 (8) | 1.2 (0.9, 1.7) | 46 (7) | 1.0 (0.8, 1.4) | |
| > 51 (116.0) | 1,321 (7) | 34 (8) | 1.1 (0.8, 1.6) | 46 (9) | 1.4 (1.0, 1.9) | 43 (7) | 0.9 (0.7, 1.3) | 0.24 |
| Missing | 1,012 | 26 | | 29 | | 47 | | |
| Trend (47.3) ^h | | | 1.1 (0.9, 1.2) | | 1.1 (1.0, 1.3) | | 1.0 (0.8, 1.1) | 0.20 |
| Malathion ^f | | | | | | | | |
| 0 (0.0) | 3,559 (34) | 77 (26) | Reference | 74 (25) | Reference | 109 (31) | Reference | |
| 1-9 (8.8) | 2,871 (27) | 74 (25) | 1.1 (0.8, 1.6) | 71 (24) | 1.2 (0.8, 1.7) | 102 (29) | 1.0 (0.8, 1.4) | |
| 10-39 (24.5) | 2,253 (21) | 65 (22) | 1.3 (0.9, 1.8) | 79 (27) | 1.6 (1.2, 2.3) | 73 (21) | 1.0 (0.7, 1.4) | |
| > 39 (108.5) | 1,939 (18) | 77 (26) | 1.8 (1.3, 2.5) | 72 (24) | 1.9 (1.4, 2.7) | 71 (20) | 1.2 (0.8, 1.6) | 0.21 |
| Missing | 368 | 13 | | 19 | | 16 | | |
| Trend (47.3) ^h | | | 1.3 (1.1, 1.4) | | 1.3 (1.1, 1.4) | | 1.1 (0.9, 1.2) | 0.07 |
| Parathion ^f | | | | | | | | |
| 0 (0.0) | 9,784 (92) | 257 (88) | Reference | 262 (89) | Reference | 336 (93) | Reference | |
| 1-9 (8.8) | 362 (3) | 16 (6) | 1.5 (0.9, 2.8) | 12 (4) | 0.8 (0.4, 1.7) | 15 (4) | 1.3 (0.7, 2.3) | |
| 10-51 (24.5) | 229 (2) | 8 (3) | 1.5 (0.6, 3.4) | 8 (3) | 1.0 (0.5, 2.1) | 6 (2) | 0.6 (0.3, 1.5) | |
| > 51 (116.0) | 236 (2) | 10 (3) | 1.4 (0.7, 2.8) | 13 (4) | 2.3 (1.3, 4.2) | 5 (1) | 0.8 (0.3, 1.9) | 0.24 |
| Missing | 379 | 15 | | 20 | | 9 | | |
| Trend (53.8) ^h | | | 1.2 (0.9, 1.6) | | 1.5 (1.1, 1.9) | | 0.9 (0.6, 1.4) | 0.13 |
| Phorate ^f | | | | | | | | |
| 0 (0.0) | 7,258 (68) | 167 (57) | Reference | 189 (64) | Reference | 259 (71) | Reference | |
| 1-9 (8.8) | 1,452 (14) | 42 (14) | 1.4 (0.9, 2.0) | 49 (17) | 1.1 (0.8, 1.6) | 41 (11) | 0.7 (0.5, 1.0) | |
| 10-25 (24.5) | 961 (9) | 39 (13) | 1.6 (1.1, 2.3) | 33 (11) | 1.3 (0.9, 1.8) | 27 (7) | 0.7 (0.5, 1.1) | |
| > 25 (62.5) | 990 (9) | 44 (15) | 1.8 (1.3, 2.6) | 25 (8) | 0.9 (0.6, 1.4) | 36 (10) | 1.0 (0.7, 1.4) | < 0.01 |

| Cumulative lifetime days personally mixed or applied ^{a,b} | Non-cases [n (%)] | PRE-E ^c Cases [n (%)] | PRE-E ^c IP-weighted OR ^d (95% CI) | PRE-B ^c Cases [n (%)] | PRE-B ^c IP-weighted OR ^d (95% CI) | POST ^c Cases [n (%)] | POST ^c IP-weighted OR ^d (95% CI) | p for difference among ORs ^e |
|---|----------------------|--|---|--|---|---------------------------------------|--|---|
| Missing | 329 | 14 | | 19 | | 8 | | |
| Trend (47.3) ^h | | | 1.6 (1.2, 2.0) | | 1.0 (0.7, 1.4) | | 0.9 (0.7, 1.2) | 0.01 |
| <i>Terbufos</i> | | | | | | | | |
| 0 (0.0) | 10,867 (59) | 227 (51) | Reference | 274 (54) | Reference | 385 (60) | Reference | |
| 1-20 (8.8) | 2,755 (15) | 73 (16) | 1.3 (1.0, 1.7) | 89 (18) | 1.2 (0.9, 1.6) | 92 (14) | 1.0 (0.8, 1.2) | |
| 21-56 (38.8) | 2,639 (14) | 75 (17) | 1.4 (1.0, 1.8) | 77 (15) | 1.1 (0.9, 1.5) | 97 (15) | 1.0 (0.8, 1.3) | |
| > 56 (116.0) | 2,208 (12) | 71 (16) | 1.5 (1.1, 1.9) | 65 (13) | 1.2 (0.9, 1.6) | 69 (11) | 0.9 (0.7, 1.2) | 0.19 |
| Missing | 1,037 | 28 | | 35 | | 45 | | |
| Trend (88.5) ^h | | | 1.3 (1.1, 1.6) | | 1.1 (0.9, 1.4) | | 0.9 (0.8, 1.2) | 0.07 |
| <i>Pyrethroid insecticides</i> | | | | | | | | |
| Permethrin (for animals) | | | | | | | | |
| 0 (0.0) | 15,726 (85) | 372 (83) | Reference | 420 (83) | Reference | 546 (84) | Reference | |
| 1-9 (8.8) | 1,195 (6) | 25 (6) | 0.9 (0.6, 1.4) | 37 (7) | 1.0 (0.7, 1.5) | 37 (6) | 1.0 (0.7, 1.5) | |
| 10-51 (24.5) | 868 (5) | 18 (4) | 0.8 (0.5, 1.4) | 29 (6) | 1.3 (0.9, 1.9) | 30 (5) | 1.0 (0.7, 1.4) | |
| > 51 (116.0) | 737 (4) | 33 (7) | 1.8 (1.2, 2.6) | 21 (4) | 1.0 (0.7, 1.6) | 34 (5) | 1.4 (1.0, 1.9) | 0.43 |
| Missing | 980 | 26 | | 33 | | 41 | | |
| Trend (47.3) ^h | | | 1.3 (1.1, 1.5) | | 1.0 (0.9, 1.2) | | 1.1 (1.0, 1.3) | 0.26 |
| Permethrin (for crops) | | | | | | | | |
| 0 (0.0) | 15,902 (87) | 377 (85) | Reference | 415 (84) | Reference | 560 (88) | Reference | |
| 1-9 (8.8) | 1,294 (7) | 37 (8) | 1.3 (0.9, 1.8) | 48 (10) | 1.5 (1.1, 2.1) | 46 (7) | 1.0 (0.7, 1.4) | |
| 10-30 (24.5) | 481 (3) | 14 (3) | 1.3 (0.8, 2.4) | 15 (3) | 1.1 (0.7, 2.0) | 5 (1) | 0.3 (0.1, 0.7) | |
| > 30 (108.5) | 685 (4) | 15 (3) | 0.9 (0.5, 1.6) | 15 (3) | 0.9 (0.5, 1.5) | 27 (4) | 1.2 (0.8, 1.8) | 0.04 |
| Missing | 1,144 | 31 | | 47 | | 50 | | |
| Trend (47.3) ^h | | | 1.0 (0.8, 1.2) | | 1.0 (0.8, 1.2) | | 1.0 (0.9, 1.3) | 0.88 |

Abbreviations: 2,4-D, (2,4-dichlorophenoxy)acetic acid; 2,4,5-T, (2,4,5-trichlorophenoxy)acetic acid; 2,4,5-TP, (RS)-2-(2,4,5-trichlorophenoxy)propionic acid; CI, confidence interval; DDT, 1,1,1-trichloro-2,2-bis(4-chlorophenyl)ethane; EPTC, S-ethyl dipropyl(thiocarbamate); IQR, interquartile range; IP, inverse probability; OR, odds ratio; POST, post-enrollment; PRE-B, pre-enrollment both; PRE-E, pre-enrollment enrollment only.

^aCategory boundaries set at tertiles of cumulative lifetime days of use for each pesticide among all male private pesticide applicators who used it.

^bFewer than five PRE-E, PRE-B, or POST cases used trichlorfon or ziram at every exposure level. ^cSee footnote a at the bottom of Table S1 for a description of the three case groups. ^dWeights adjusted for age at enrollment (modeled with a restricted, quadratic spline with knots at ages 40, 48, and 57 years based on percentiles of the age distribution in cases), ever diagnosed with diabetes, education level, state of residence, not missing covariate data (conditional on age, state, the exposure, and pairwise interaction terms between each covariate and the exposure), and not dropping out of the AHS cohort (conditional on age, diabetes, education, state, the exposure, and pairwise interaction terms between each covariate and the exposure). 95% CIs calculated with robust variance estimates. ^eDifferences among case-group-specific ORs tested via Wald χ^2 tests. ^fData available only for 11,982 applicators who completed the farmer questionnaire. Weights additionally adjusted for completing the farmer questionnaire (conditional on age, diabetes, education, and state). ^gCategory boundaries set at the median of cumulative lifetime days of use for each pesticide among all male private pesticide applicators who used it. ^hUsed within-category medians and scaled the OR to an IQR-unit (days) increase in cumulative lifetime days of use for each pesticide among all male private pesticide applicators who used it. ⁱOR (95% CI) not shown because fewer than five PRE-E or PRE-B cases used aldicarb, captan, carbon tetrachloride/carbon disulfide (80/20 mix), chlordane, dieldrin, ethylene dibromide, maneb/mancozeb, or paraquat at this exposure level. ^jBenomyl is also included in carbamates.

Table S6. Pesticide use and self-reported depression among male private pesticide applicators in the Agricultural Health Study without weighting for potential biases from missing covariate data, missing farmer questionnaire, or drop out.

| Variable | PRE-E ^a Adjusted OR ^b (95% CI) | PRE-B ^a Adjusted OR ^b (95% CI) | POST ^a Adjusted OR ^b (95% CI) | p for Difference among ORs ^c |
|---|--|--|---|---|
| Cumulative days personally mixed or applied pesticides ^d | | | | |
| ≤ 56 (Median = 24.5) | Reference | Reference | Reference | |
| 57-225 (116.0) | 1.3 (1.0, 1.7) | 1.1 (0.9, 1.4) | 0.9 (0.7, 1.1) | |
| 226-457 (369.8) | 1.4 (1.0, 1.8) | 1.2 (0.9, 1.6) | 1.1 (0.9, 1.4) | |
| > 457 (767.3) | 1.5 (1.1, 2.0) | 1.1 (0.8, 1.5) | 1.0 (0.8, 1.2) | 0.29 |
| Missing | | | | |
| Trend (IQR = 401.3) ^e | 1.2 (1.0, 1.3) | 1.0 (0.9, 1.2) | 1.0 (0.9, 1.2) | 0.26 |
| Ever diagnosed with pesticide poisoning ^f | | | | |
| No | Reference | Reference | Reference | |
| Yes | 5.1 (3.4, 7.7) | 2.7 (1.6, 4.5) | 1.0 (0.5, 2.2) | < 0.01 |
| Missing | | | | |
| Ever experienced an incident of unusually high personal pesticide exposure ^f | | | | |
| No | Reference | Reference | Reference | |
| Yes | 2.3 (1.8, 3.0) | 2.2 (1.7, 2.8) | 1.1 (0.8, 1.4) | < 0.01 |
| Missing | | | | |

Abbreviations: CI, confidence interval; IQR, interquartile range; OR, odds ratio; POST, post-enrollment; PRE-B, pre-enrollment both; PRE-E, pre-enrollment enrollment only.

^aSee footnote a at the bottom of Table S1 for a description of the three case groups. ^bAdjusted for age at enrollment (modeled with a cubic polynomial), ever diagnosed with diabetes, education level, and state of residence. ^cDifferences among case-group-specific ORs tested via Wald χ^2 tests. ^dCategory boundaries set at quartiles of cumulative days of pesticide use among all male private pesticide applicators. ^eUsed within-category medians and scaled the OR to an IQR-unit (days) increase in cumulative days of pesticide use among all male private pesticide applicators.

^fData available only for 11,982 applicators who completed the farmer questionnaire.

Table S7. Ever-use of pesticide classes and specific pesticides and self-reported depression among male private pesticide applicators in the Agricultural Health Study without weighting for potential biases from missing covariate data, missing farmer questionnaire, or drop out.

| Ever personally mixed or applied | PRE-E ^a Adjusted OR ^{b,c} (95% CI) | PRE-B ^a Adjusted OR ^{b,c} (95% CI) | POST ^a Adjusted OR ^{b,c} (95% CI) | p for Difference among ORs ^d |
|---|--|--|---|---|
| Fumigants | 1.3 (1.1, 1.7) | 1.7 (1.4, 2.1) | 1.2 (1.0, 1.5) | 0.04 |
| Aluminum phosphide | 1.5 (1.0, 2.1) | 1.5 (1.1, 2.1) | 1.5 (1.1, 2.0) | 1.00 |
| Carbon tetrachloride/ carbon disulfide (80/20 mix) | 1.5 (1.1, 2.0) | 1.6 (1.2, 2.2) | 1.2 (0.9, 1.7) | 0.40 |
| Ethylene dibromide | 1.5 (1.0, 2.2) | 1.4 (0.9, 2.1) | 1.2 (0.8, 1.8) | 0.83 |
| Methyl bromide | 1.2 (0.9, 1.6) | 1.4 (1.0, 1.8) | 1.0 (0.8, 1.3) | 0.31 |
| Fungicides | 1.2 (1.0, 1.5) | 1.4 (1.1, 1.7) | 1.1 (0.9, 1.3) | 0.24 |
| Benomyl ^e | 1.1 (0.8, 1.6) | 1.0 (0.7, 1.4) | 1.1 (0.8, 1.4) | 0.82 |
| Captan | 1.1 (0.9, 1.5) | 1.4 (1.1, 1.8) | 1.1 (0.9, 1.4) | 0.29 |
| Chlorothalonil | 1.0 (0.7, 1.5) | 1.4 (1.0, 1.9) | 1.1 (0.8, 1.5) | 0.49 |
| Maneb/mancozeb | 1.2 (0.9, 1.7) | 1.1 (0.8, 1.5) | 1.0 (0.8, 1.3) | 0.70 |
| Metalaxyl | 1.4 (1.1, 1.8) | 1.2 (1.0, 1.5) | 1.0 (0.8, 1.2) | 0.09 |
| Ziram | 1.5 (0.8, 2.9) | 0.7 (0.3, 1.6) | 1.2 (0.7, 2.2) | 0.32 |
| Herbicides | 1.7 (0.7, 4.2) | 1.3 (0.6, 2.8) | 1.4 (0.7, 2.6) | 0.90 |
| Alachlor | 1.2 (1.0, 1.5) | 1.2 (1.0, 1.4) | 1.1 (1.0, 1.3) | 0.81 |
| Butylate | 1.0 (0.8, 1.2) | 1.1 (0.9, 1.3) | 1.1 (0.9, 1.3) | 0.72 |
| Chlorimuron-ethyl | 1.0 (0.8, 1.2) | 1.1 (0.9, 1.3) | 1.1 (0.9, 1.3) | 0.63 |
| Dicamba | 0.9 (0.7, 1.1) | 1.0 (0.8, 1.3) | 1.1 (0.9, 1.3) | 0.47 |
| EPTC | 1.2 (1.0, 1.5) | 0.9 (0.7, 1.1) | 1.2 (1.0, 1.4) | 0.13 |
| Glyphosate | 1.2 (1.0, 1.5) | 1.1 (0.9, 1.4) | 1.1 (0.9, 1.3) | 0.72 |
| Imazethapyr | 1.1 (0.9, 1.3) | 0.9 (0.7, 1.0) | 1.1 (0.9, 1.3) | 0.21 |
| Metolachlor | 1.1 (0.9, 1.3) | 0.8 (0.7, 1.0) | 1.0 (0.8, 1.1) | 0.14 |
| Paraquat | 1.2 (0.9, 1.5) | 1.1 (0.9, 1.4) | 1.0 (0.8, 1.2) | 0.67 |
| Pendimethalin | 1.2 (1.0, 1.4) | 0.9 (0.8, 1.1) | 0.9 (0.8, 1.1) | 0.10 |
| Petroleum oil | 1.3 (1.1, 1.6) | 1.2 (1.0, 1.5) | 1.0 (0.9, 1.2) | 0.18 |
| Trifluralin | 1.2 (1.0, 1.4) | 1.1 (0.9, 1.3) | 1.1 (0.9, 1.3) | 0.73 |
| Phenoxy herbicides | 1.1 (0.8, 1.4) | 1.2 (0.9, 1.6) | 0.9 (0.8, 1.1) | 0.24 |
| 2,4-D | 1.0 (0.8, 1.3) | 1.1 (0.9, 1.4) | 0.9 (0.8, 1.1) | 0.57 |
| 2,4,5-T | 1.4 (1.2, 1.8) | 1.5 (1.2, 1.8) | 1.1 (0.9, 1.4) | 0.09 |
| 2,4,5-TP | 1.5 (1.1, 1.9) | 1.4 (1.1, 1.8) | 1.1 (0.8, 1.4) | 0.26 |
| Triazine herbicides | 1.1 (0.8, 1.4) | 1.0 (0.8, 1.2) | 1.0 (0.8, 1.3) | 0.82 |
| Atrazine | 1.2 (0.9, 1.5) | 1.0 (0.8, 1.3) | 1.0 (0.8, 1.2) | 0.54 |
| Cyanazine | 1.3 (1.0, 1.6) | 1.1 (0.9, 1.4) | 1.1 (0.9, 1.3) | 0.63 |
| Metribuzin | 1.1 (0.9, 1.3) | 1.0 (0.9, 1.3) | 1.0 (0.9, 1.2) | 0.91 |

| Ever personally mixed or applied | PRE-E^a Adjusted OR^{b,c} (95% CI) | PRE-B^a Adjusted OR^{b,c} (95% CI) | POST^a Adjusted OR^{b,c} (95% CI) | p for Difference among ORs^d |
|---|---|---|--|---|
| Insecticides | 1.4 (0.8, 2.4) | 0.9 (0.6, 1.4) | 1.5 (1.0, 2.4) | 0.20 |
| Carbamates ^e | 1.1 (0.9, 1.3) | 1.2 (1.0, 1.5) | 1.1 (0.9, 1.4) | 0.68 |
| Aldicarb | 0.9 (0.7, 1.3) | 1.1 (0.8, 1.5) | 1.2 (0.9, 1.6) | 0.45 |
| Carbaryl | 1.2 (1.0, 1.5) | 1.3 (1.1, 1.6) | 1.2 (1.0, 1.4) | 0.75 |
| Carbofuran | 1.1 (0.9, 1.3) | 1.2 (1.0, 1.4) | 0.9 (0.8, 1.1) | 0.16 |
| Organochlorine insecticides | 1.9 (1.5, 2.3) | 1.2 (1.0, 1.5) | 1.2 (1.0, 1.5) | 0.01 |
| Aldrin | 1.4 (1.1, 1.7) | 1.5 (1.2, 1.8) | 1.2 (1.0, 1.5) | 0.40 |
| Chlordane | 1.5 (1.3, 1.9) | 1.2 (1.0, 1.5) | 1.1 (0.9, 1.3) | 0.05 |
| DDT | 1.3 (1.1, 1.7) | 1.2 (0.9, 1.4) | 0.9 (0.7, 1.1) | 0.03 |
| Dieldrin | 1.4 (1.0, 1.9) | 1.3 (1.0, 1.8) | 1.1 (0.8, 1.5) | 0.60 |
| Heptachlor | 1.6 (1.3, 2.0) | 1.3 (1.0, 1.6) | 1.0 (0.8, 1.2) | 0.01 |
| Lindane | 1.6 (1.3, 1.9) | 1.3 (1.0, 1.5) | 1.1 (0.9, 1.4) | 0.05 |
| Toxaphene | 1.3 (1.0, 1.6) | 1.4 (1.1, 1.7) | 1.1 (0.9, 1.4) | 0.49 |
| Organophosphate insecticides | 1.5 (1.0, 2.1) | 1.1 (0.8, 1.5) | 1.3 (1.0, 1.8) | 0.51 |
| Chlorpyrifos | 1.2 (1.0, 1.4) | 1.3 (1.1, 1.6) | 1.0 (0.9, 1.2) | 0.07 |
| Coumaphos | 1.3 (1.0, 1.7) | 1.3 (1.0, 1.6) | 0.9 (0.7, 1.1) | 0.08 |
| Diazinon | 1.3 (1.1, 1.6) | 1.4 (1.1, 1.6) | 1.1 (1.0, 1.4) | 0.39 |
| Dichlorvos | 1.0 (0.8, 1.4) | 1.5 (1.2, 1.9) | 1.3 (1.1, 1.6) | 0.10 |
| Fonofos | 1.3 (1.0, 1.6) | 1.2 (0.9, 1.4) | 1.0 (0.8, 1.2) | 0.17 |
| Malathion | 1.4 (1.1, 1.7) | 1.2 (1.0, 1.5) | 1.2 (1.0, 1.4) | 0.56 |
| Parathion | 1.5 (1.2, 1.9) | 1.2 (0.9, 1.5) | 1.2 (1.0, 1.5) | 0.30 |
| Phorate | 1.3 (1.0, 1.6) | 1.0 (0.9, 1.3) | 1.0 (0.9, 1.2) | 0.22 |
| Terbufos | 1.4 (1.1, 1.7) | 1.2 (1.0, 1.4) | 1.0 (0.8, 1.2) | 0.02 |
| Trichlorfon | 1.7 (0.7, 4.1) | f | f | f |
| Pyrethroid insecticides | 1.2 (1.0, 1.5) | 1.2 (1.0, 1.4) | 0.9 (0.8, 1.1) | 0.09 |
| Permethrin (for animals) | 1.2 (1.0, 1.6) | 1.1 (0.9, 1.4) | 1.0 (0.8, 1.3) | 0.48 |
| Permethrin (for crops) | 1.2 (0.9, 1.6) | 1.3 (1.0, 1.7) | 0.9 (0.7, 1.1) | 0.04 |

Abbreviations: 2,4-D, (2,4-dichlorophenoxy)acetic acid; 2,4,5-T, (2,4,5-trichlorophenoxy)acetic acid; 2,4,5-TP, (RS)-2-(2,4,5-trichlorophenoxy)propionic acid; CI, confidence interval; DDT, 1,1,1-trichloro-2,2-bis(4-chlorophenyl)ethane; EPTC, S-ethyl dipropyl(thiocarbamate); OR, odds ratio; POST, post-enrollment; PRE-B, pre-enrollment both; PRE-E, pre-enrollment enrollment only.

^aSee footnote a at the bottom of Table S1 for a description of the three case groups. ^bMale private pesticide applicators who did not use each pesticide class or specific pesticide were the reference.

^cAdjusted for age at enrollment (modeled with a cubic polynomial), ever diagnosed with diabetes, education level, and state of residence. ^dDifferences among case-group-specific ORs tested via Wald χ^2 tests. ^eBenomyl is also included in carbamates. ^fOR (95% CI) and p for difference not shown because fewer than five PRE-B or POST cases ever personally mixed or applied trichlorfon.

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